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1. search the following applicants

A. C. Richard Schlegel

B. A Bennett Jensen

search APS, DERWENT, and literature references.

2. search the following keywords

A) papillomavirus

HPV?

B) capsid

C) L1 protein

search APS, DERWENT and literature references

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Number of Databases:	Structure	D
	Bibliographic	

VIROLOGY

Volume 187, Number 2, April 1992

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b350,351,357,358

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File 350:Derwent World Patents Index

1963-1980, EQUIVALENTS THRU DW=9247

**FILE350: Format 9 includes the expanded patent table. Preformatted REPORTS are available. Type ?FMT350, ?NEWS350, ?RATES350 for more info.

File 351:DERWENT WORLD PATENTS INDEX-LATEST

1981+;DW=9301,UA=9241,UM=9214

**FILE351: Format 9 includes the expanded patent table. Preformatted REPORTS are available. Type ?FMT351, ?NEWS351, ?RATES351 for more info.

File 357:DERWENT BIOTECHNOLOGY ABS 1982-1993/FEB

(Copr. 1993 Derwent Pub. ltd.)

File 358:CURRENT BIOTECHNOLOGY ABS 1983-1993/MAR

(COPR. 1993 ROYAL SOC CHEM)

Set	Items	Description
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?s	papillomavir?	or hpv or pv) and capsid and (l1 protein or l1)	
		83	PAPILLOMAVIR?
		146	HPV
		1447	PV
		462	CAPSID
		0	L1 PROTEIN
S1	4	L1	
		(PAPILLOMAVIR? OR HPV OR PV) AND CAPSID AND (L1 PROTEIN OR L1)	

?t1/5/1-4;e auhshgael c

1/5/1 (Item 1 from file: 351)

004775488 WPI Acc No: 86-278829/42

XRAM Acc No: C86-120579

Type-specific papillomavirus DNA sequences and peptide(s) - useful in assays for specific Papillomavirus and in vaccines; DEOXYRIBONUCLEIC ACID

Patent Assignee: (GEOU) UNIV GEORGETOWN; (GEOU) GEORGETOWN UNIV

Author (Inventor): JENSON A B; LANCASTER D W; JENSON B A; LANCASTER W D

Number of Patents: 006

Number of Countries: 014

Patent Family:

CC Number	Kind	Date	Week	
WO 8605816	A	861009	8642	(Basic)
EP 217919	A	870415	8715	
JP 62502378	W	870917	8743	
US 5057411	A	911015	9144	
EP 217919	B1	920805	9232	
DE 3686304	G	920910	9238	

Applications (CC,No,Date): DE 3686304 (860328); EP 8602614 (860328); WO 86US629 (860328); WO 86US629 (860328); JS629 (860328); EP 8690261 (860328); JP 86502314 (860328); US 346283 (890501); EP 86902614 (860328); WO 86US629 (860328)

Language: English

EP and/or WO Cited Patents: 2.Jnl.Ref; US 4551270; 3.Jnl.Ref; EP 133123; EP 192001; EP 92456; US 4358535; US 4419446

Designated States

(National): DK; JP; SE

(Regional): AT; BE; CH; DE; FR; GB; IT; LU; NL; LI; SE

Filing Details: DE3686304 Based on EP 217919; DE3686304 Based on WO 8605816; EP0217919 Based on WO 8605816

Abstract (Basic): WO 8605816

The following are claimed: (1) a detectably labelled polynucleotide sequence specific for a given papilloma-virus (PV) type, pref. of 15-75 nucleotides; (2) isolated polynucleotide segment comprising a sequence coding for a type-specific PV gene prod., pref. of 15-75 nucleotides; (3) a polypeptide (I) having a sequence of aminoacids specific for a particular papillomavirus and representing a relatively small fragment of the naturally occurring L1 open reading frame polypeptide; (4) a method for identifying type-specific PV comprising contacting a sample contg. the virus with labelled antibody which is immunologically specific for the virus and determining the extent of binding of the antibody to the virus; and (5) a polypeptide fragment (II) of the L1 capsid protein having a genus-specific sequence of aminoacids.

More specifically (I) is formula: KNNKGDATLK. (II) is of formula RGQPLG.

USE/ADVANTAGE - Assays for type-specific PV, including DNA probes, RNA probes and immunoassays are now possible. Vaccines against specific PV's may also be produced. Thus specific etiology of papilloma and closely associated carcinoma can be identified. @(60pp Dwg.No.0/6)@

Abstract (US): 9144 US 5057411

New detectably-labelled polynucleotide sequence comprises a papillomavirus (PV) polynucleotide sequence to distinguish between PV types. Sequence has at least 15 (pref. 18-75) nucleotides of an L1 open reading frame type-specific sequence, but less than the entire genome. The polynucleotide segments are isolated.

Characterising PV type comprises contacting a sample contg. single stranded PV DNA with a PV type-specific polynucleotide probe specific for the PV type, and hybridising between the DNA and the probe. The probe-DNA hybrids are detected, the PV DNA characterised by presence or absence of hybrids. The probe is labelled e.g. by radio, fluorescence, chemiluminescent, enzyme, antibody, or free radical label. The probe is e.g. specific labelled DNA or RNA probe.

USE - By identifying segments common to all PV's and others confined to each individual PV, identification of individual (type-specific) or genus-specific PV is possible. @(22pp)@

Abstract (EP): 9232 EP 217919 B

A detectably labelled polynucleotide segment for distinguishing between papillomavirus types, said segment comprising at least 5 nucleotides of the nucleotide sequence of a given papillomavirus type which corresponds to BPV-1 L1 protein amino acids 251-291. Dwg.0/6

File Segment: CPI

Derwent Class: B04; D16;

Int Pat Class: A61K-039/42; C07H-015/12; C07K-007/06; C12N-015/37; C12Q-001/68

Manual Codes (CPI/A-N): B04-B02B4; B04-B03B; B04-B04A1; B04-B04C6; B04-C01B; B11-C07A; B12-K04A1; B12-K04A4; D05-H06; D05-H12

Chemical Fragment Codes (M1):

01 M423 M760 M903 N102 N133 N134 N135 N136 Q233 V600 V644 V754

02 M423 M750 M903 N102 N133 N134 N135 N136 Q233 V500 V560 V753

03 C811 M423 M430 M710 M781 M782 M903 N102 N133 N134 N135 Q233 Q505

V600 V611 V753 V802 V810

05 C811 H1 H100 H181 H182 H4 H401 H481 H8 J011 J012 J1 J171
 J172 J3 J371 M280 M311 M312 M313 M315 M321 M331 M332 M333 M342 M343
 M349 M381 M391 M421 M423 M430 M510 M520 M530 M540 M620 M710 M781 M782
 M903 N102 N133 N134 N135 P210 P831 Q233 Q505 V279 V600 V611 V802 V810
 V901 V912 V913 V921
 06 C811 F012 F423 H1 H100 H181 J0 J011 J012 J1 J111 J171 J3 J371 K0
 L2 L250 M280 M311 M313 M315 M320 M321 M332 M333 M342 M343 M349 M381
 M391 M423 M430 M510 M520 M521 M530 M540 M620 M710 M781 M782 M903 N102
 N133 N134 N135 P831 Q233 Q505 V600 V611 V802 V810 V901 V912 V921
 Chemical Fragment Codes (M6):
 07 M903 P210 P831 Q233 Q505 R309 R513 R514 R515 R521 R614 R621 R623
 R624 R625 R626 R627 R635 R639

1/5/2 (Item 1 from file: 357)

128888 DBA Accession No.: 92-01380

Expression of vaccinia recombinant HPV 16 L1 and L2 ORF proteins in
 epithelial cells is sufficient for assembly of HPV virion-like
 particles - human papilloma virus recombinant L1 and L2 protein
 production; vaccinia virus vector construction; virus-like particle
 expression in CV-1 cell culture; potential recombinant vaccine

AUTHOR: Zhou J; Sun X Y; Stenzel D J; +Frazer I H

CORPORATE SOURCE: Lions Human Immunology Laboratory, Princess Alexandra
 Hospital, Brisbane, Queensland 4102, Australia.

JOURNAL: Virology (185, 1, 251-57) 1991 CODEN: VIRLAX

LANGUAGE: English

ABSTRACT: Vaccinia virus (VV) plasmid pLC201VV was designed to co-express
 the L1 and L2 late genes of human papilloma virus type-16 (HPV16). The
 L1 gene from plasmid pHPV16 was cloned under the VV 4b promoter to form
 plasmid pLC200. The L2 gene from plasmid pHPV16 was cloned under the
 control of the VV 28K late promoter, and then cloned into plasmid
 pLC200 to form plasmid pLC201. An EcoRI-XbaI DNA fragment containing
 the E1/E4 gene from W12 cell cDNA and under the control of the VV 11K
 promoter was cloned into plasmid pLC201 to form plasmid pLC202. pLC201
 and pLC202 were used to construct VV recombinants (pLC201VV and
 pLC202VV). L1 and L2 production occurred in CV-1 cells infected with
 pLC201VV, and 40 nm virus-like particles (VLPs) of density 1.31 g/ml
 were produced in the nuclei of cells producing both L1 and L2, but not
 in cells producing either L1 or L2. VLPs were isolated from cells by
 sucrose gradient sedimentation and shown to consist of capsomeres
 similar to HPV and contain glycosylated L1 viral capsid protein. The VV
 production method for HPV VLPs may be useful for biochemical studies
 and recombinant vaccine construction. (38 ref)

DESCRIPTORS: human papilloma virus recombinant L1 protein prep., L2 protein
 prep., late gene cloning, co-expression, vaccinia virus vector
 construction, virus-like particle expression in CV-1 cell culture, pot.
 recombinant vaccine mammal monkey kidney

SECTION: Pharmaceuticals-Vaccines; Microbiology-Genetics; Cell Culture-
 Animal Cell Culture (D4,A1,J1)

1/5/3 (Item 2 from file: 357)

122167 DBA Accession No.: 91-09809

Expression of human papilloma virus proteins in yeast *Saccharomyces*
cerevisiae - protein secretion as fusion protein with yeast
 prepro-alpha-factor

AUTHOR: Carter J J; Yaegashi N; Jenison S A; +Galloway D A

CORPORATE SOURCE: Fred Hutchinson Cancer Research Center, Seattle,
 Washington 98104, USA.

JOURNAL: Virology (182, 2, 513-21) 1991 CODEN: VIRLAX

LANGUAGE: English

ABSTRACT: The L1 and L2 proteins of human papilloma virus (HPV) types 1, 6
 and 16, and the E7 proteins of HPV 16 were expressed in *Saccharomyces*
cerevisiae. The yeast-expressed proteins were readily detected by
 immunoblotting and were generally intact. The recombinant HPV 1 L2 and
 L2 proteins were indistinguishable from the major and minor capsid

proteins were secreted from yeast by fusion to the prepro-alpha-factor signal peptide. Following secretion of the HPV-16 E7 protein, a rapid method of purification was developed. These recombinant proteins were of the mol.wt. expected for the major and minor virion proteins. The yeast-expressed proteins were used as antigens to study the human immune response in Western blot assays, ELISA and in immune precipitation. One human serum reacted with intact, but not denatured HPV-16 L2 proteins, suggesting that the yeast-expressed proteins will be useful to detect antibodies reactive with conformational epitopes. (34 ref)

DESCRIPTORS: human papilloma virus recombinant protein expression in *Saccharomyces cerevisiae*, protein secretion as fusion protein with yeast prepro-alpha-factor mammal fungus

SECTION: Pharmaceuticals-Other; Microbiology-Genetics (D5,A1)

1/5/4 (Item 3 from file: 357)

071415 DBA Accession No.: 88-01763

Expression of human papilloma virus type 6 and type 16 capsid proteins in bacteria and their antigenic characterization - *Escherichia coli* expression of beta-galactosidase fusion proteins; vaccine and diagnostic reagent development

AUTHOR: Banks L; Matlashewski G; Pim D; Churcher M; Roberts C; Crawford L

CORPORATE AFFILIATE: Wellcome

CORPORATE SOURCE: Department of Biochemical Virology, The Wellcome Research Laboratories, Langley Court, Beckenham, Kent, BR3 3BS, UK.

JOURNAL: J.Gen.Virol. (68, Pt.12, 3081-89) 1987 CODEN: JGVIAY

LANGUAGE: English

ABSTRACT: The L1 and L2 capsid proteins encoded by human papilloma virus types 6 and 16 (HPV-6 and HPV-16) have been produced in *Escherichia coli*. Antisera were raised against the HPV-6 L1- and L2-beta-galactosidase (EC-3.2.1.23) fusion proteins and against a HPV-16 L1 C-terminal peptide 14 amino acids long. The HPV-16 L1 peptide antibodies are highly reactive with the HPV-16 L1-beta-galactosidase fusion protein but not against the equivalent HPV-6 L1-beta-galactosidase fusion protein. The effectiveness of these antibodies was compared with commercially available anti-bovine papilloma virus type 1 (BPV-1) antibodies. The anti-BPV-1 antibodies reacted well against HPV-6 L1-beta-galactosidase but not against HPV-16 L1-beta-galactosidase. The L2 portion of the HPV-6 L2-beta-galactosidase fusion protein was particularly immunogenic, since antibodies raised against it were predominantly reactive with the L2 moiety. The HPV-16 L1 peptide antibodies described will be preferred reagents for the specific detection of HPV-16 capsid antigens, which may be particularly important in early diagnosis of HPV-16 infection. (22 ref)

E.C. NUMBERS: 3.2.1.23

DESCRIPTORS: human papilloma virus type 6, type 16 capsid protein, beta-galactosidase fusion protein etc. expression in *Escherichia coli*, pot. vaccine, diagnostic reagent development bacterium mammal enzyme EC-3.2.1.23

SECTION: Pharmaceuticals-Vaccines; Pharmaceuticals-Other; Microbiology-Genetics (D4,D5,A1)

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E2	24	*AU=SCHLEGEL C
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1136 PAPILOMAVIR?/AB
1322 PAPILOMAVIR?/IA
 (PAPILOMAVIR?/BI,AB)
1554 CAPSID/BI
2930 CAPSID/AB
3348 CAPSID/IA
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1143 "L1"/BI
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L7 10 (PAPILOMAVIR? AND CAPSID AND "L1 PROTEIN")/IA

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L7 ANSWER 1 OF 10 COPYRIGHT 1993 ACS
TI Self-assembly of human papillomavirus type 1 capsids by
 expression of the L1 protein along or by
 coexpression of the L1 and L2 ***capsid*** proteins
SO J. Virol., 67(1), 315-22
AU Hagensee, Michael E.; Yaegashi, Nobuo; Galloway, Denise A.
PY 1993
AN CA118(7):55781r
AB Vaccinia virus vectors were used to express the major (L1) and (L2)
 capsid proteins of human papillomavirus type 1
 (HPV-1) with the vaccinia virus early (p7.5K) or late (pSynth, p11K)
 promoters. All constructs expressed the appropriate-sized HPV

proteins, and both L1 and L2, singly or in combination, localized to the nucleus. Capsids were purified by cesium chloride density gradient centrifugation from nuclei of cells infected with a vaccinia virus-L1 (vac-L1) recombinant or a vac-L1-L2 recombinant but not from vac-L2-infected cells. Electron microscopy showed that the particles were 55 nm in diam. and had icosahedral symmetry. Immunogold-labeled antibodies confirmed the presence of the L1 and L2 proteins in the HPV-1 capsids. Capsids contg. L1 alone were fewer and more variable in size and shape than capsids contg. the L1 and L2 proteins. The L1-plus-L2 capsids were indistinguishable in appearance from HPV-1 virions obtained from plantar warts. The ability to produce HPV capsids in vitro will be useful in many studies of HPV pathogenicity.

L7 ANSWER 2 OF 10 COPYRIGHT 1993 ACS

TI HPV-1 L1 protein expressed in cos cells displays conformational epitopes found on intact virions

SO Virology, 190(1), 548-52

AU Ghim, Shin Je; Jenson, A. Bennett; Schlegel, Richard

PY 1992

AN CA117(19):189783f

AB Seven polyclonal and monoclonal antibodies were characterized for their ability to react specifically with either conformational or nonconformational epitopes of the human papillomavirus (HPV)-1 virion. Using these antibodies, it was shown that the HPV-1 L1 protein (when expressed by an SV40 vector in cos cells) displayed conformational epitopes characteristic of intact viral particles. In addn., the L1 capsid protein was translocated normally into cell nuclei, was of appropriate size (57 kDa), and could be isolated in native form by immunopptn. techniques. Most importantly, the screening of expressed papillomavirus capsid proteins for reactivity with conformation-dependent antibodies represents a new, general methodol. for ensuring that such proteins will be suitable for use in vaccine development or in the serol. detection/typing of human papillomavirus infections.

L7 ANSWER 3 OF 10 COPYRIGHT 1993 ACS

TI Definition of linear antigenic regions of the HPV16 L1 capsid protein using synthetic virion-like particles

SO Virology, 189(2), 592-9

AU Zhou, Jian; Sun, Xiao-Yi; Davies, Huw; Crawford, Lionel; Park, David; Frazer, Ian H.

PY 1992

AN CA117(17):169102e

AB Mice of 3 haplotypes (H-2d, H-2b, and H-2d/b) were immunized with synthetic human papillomavirus (HPV)16-like particles (VLPs), produced using a vaccinia virus doubly recombinant for the L1 and L2 proteins of HPV16. The resultant anti-VLP antisera recognized HPV16 capsids by ELISA assay and baculovirus recombinant HPV16 L1 and L2 protein on immunoblot. Overlapping peptides corresponding to the HPV16 L1 amino acid sequences were used to define the immunoreactive regions of the L1 protein. The majority of the L1 peptides were reactive with IgG from the mice immunized with the synthetic HPV16 capsids. A computer algorithm predicted 7 B epitopes in HPV16 L1, 5 of which lay within peptides strongly reactive with the murine antisera. The murine anti-VLP antisera failed to react with the 2 peptides recognized by anti-HPV16L1 monoclonal antibodies raised by others against recombinant L1 fusion protein. Thus, immunoreactive epitopes of HPV16 defined using virus-like particles differ significantly from those defined using recombinant HPV16 L1 fusion proteins, which

implies that such fusion proteins may not be the antigens to look for in HPV16L1-specific immune responses in HPV-infected patients.

L7 ANSWER 4 OF 10 COPYRIGHT 1993 ACS

TI Identification of the nuclear localization signal of human papillomavirus type 16 L1 protein

SO Virology, 185(2), 625-32

AU Zhou, Jian; Doorbar, John; Sun, Xiao Yi; Crawford, Lionel V.; McLean, Cornelia S.; Frazer, Ian H.

PY 1991

AN CA116(5):38896y

AB Human papillomavirus type 16 (HPV16) L1 and L2

capsid proteins can be detected only in the nucleus of infected cells. For other nuclear proteins, specific sequences of basic amino acids (aa) termed nuclear localization signals (NLS) direct the protein from the cytoplasm to the nucleus. The authors used a series of deletion and substitution mutations of the HPV16 L1 protein, produced by recombinant vaccinia virus (rvv), to identify NLS within HPV16 L1 and showed that HPV16 L1 contains two NLS sequences, each contg. basic aa clusters. One NLS consisted of 6 basic aa (amino acids): KRKKRK from aa 525 to 530, at the carboxy terminal end of L1. The other NLS contained 2 basic aa clusters (KKR from aa 510 to 512 and KR at aa 525, 526) sepd. by 12 amino acids. Mutations in either NLS did not alter nuclear localization of L1 when the other remained intact, but mutations to both prevented nuclear localization of L1. The L1 NLS could be overridden by introduction of a membrane binding sequence at the amino terminal end of the protein. A database search showed that all sequenced papillomaviruses are predicted to have L1 and L2 capsid proteins with sequences of basic amino acids homologous with one or both NLS of HPV16 L1.

L7 ANSWER 5 OF 10 COPYRIGHT 1993 ACS

TI Characterization of murine polyclonal antisera and monoclonal antibodies generated against intact and denatured human papillomavirus type 1 virions

SO J. Virol., 65(3), 1578-83

AU Yaegashi, Nobuo; Jenison, Steven A.; Valentine, Janette M.; Dunn, Maureen; Taichman, Lorne B.; Baker, David A.; Galloway, Denise A.

PY 1991

AN CA114(17):162081b

AB Human papillomavirus type 1 (HPV1) virions, both as intact virion particles (IVP) and as detergent-denatured virions (DDV), were used to prep. polyclonal antisera and monoclonal antibodies (MAbs) in BALB/c mice. Anti-IVP antiserum contained type-specific HPV1 L2-reactive antibodies and no detectable HPV1 L1-reactive antibodies. Anti-IVP MAbs recognized a linear epitope between L2 amino acids 102 and 108 (PIDVVD). Anti-DDV antiserum contained type-specific HPV1 L1-reactive and HPV1 L2-reactive antibodies. An anti-DDV MAb recognized a linear epitope between L1 amino acids 127 and 133 (AENPTNY). HPV1a L1- and L2-encoded polypeptides expressed in *Saccharomyces cerevisiae* and by in vitro translation were equiv. in size to the major and minor virion capsid proteins, resp.

L7 ANSWER 6 OF 10 COPYRIGHT 1993 ACS

TI Human papillomavirus type 1 produces redundant as well as polycistronic mRNAs in plantar warts

SO J. Virol., 64(6), 3144-9

AU Palermo-Dilts, Deborah A.; Broker, Thomas R.; Chow, Louise T.

PY 1990

AN CA113(7):53290c

AB Human papilloma virus type 1 (HPV-1) causes plantar warts. On the basis of previously mapped mRNAs and sequence homologies of HPV-1 to other papillomaviruses, the authors designed oligonucleotide primers and employed the polymerase chain reaction to recover HPV-1 cDNAs from plantar warts. Seven spliced RNA species were characterized, including three not previously detected, and the coding potentials of each were deduced. The most abundant viral mRNA encodes an E1i E4 protein. One new species is predicted to encode the full-length E2 protein, and another can, theor., encode the E2-C or E1-M proteins, three products that regulate mRNA transcription and DNA replication. One RNA species originating from a novel HPV promoter in the upstream regulatory region has the potential to encode the minor capsid protein L2. A newly reorganized E5a open reading frame (ORF) is contained in all mRNAs that are polyadenylated at the E-region poly(A) site and also in a putative L2 mRNA. Three distinct species, two of which are derived from the upstream regulatory region promoter, have the potential to encode the L1 protein; the third species also contains the entire coding region of the E1i E4 protein 5' to the L1 ORF. Both the E1i E4 mRNA and the potentially bicistronic L1 mRNA are derived from a promoter located in the E7 ORF. The authors uncovered no evidence of alternatively spliced mRNAs that could account for the multiple, abundant E4 proteins in plantar warts, suggesting that posttranslational modification is mainly responsible for the observed protein heterogeneity.

L7 ANSWER 7 OF 10 COPYRIGHT 1993 ACS

TI Identification of immunogenic regions of the major coat protein of human papillomavirus type 16 that contain type-restricted epitopes

SO J. Gen. Virol., 70(11), 2973-87

AU Cason, John; Patel, Daksha; Naylor, Jennifer; Lunney, Declan; Shepherd, Philip S.; Best, Jennifer M.; McCance, Dennis J.
PY 1989

AN CA112(13):116820c

AB Regions were identified of the major capsid protein, L1, of the human papillomavirus (HPV) type 16 (HPV-16 L1), that are recognized by 5 monoclonal antibodies (MAbs) raised to a bacterial fusion protein contg. residues 172-375 of HPV-16 L1. All 5 MAbs recognized HPV-16-infected tissue sections by immunohistochem., but not sections infected with HPV-1a (cutaneous warts), HPV-6b or -11 (genital warts). MAbs 3D1, 5A4, and 1D6 also recognized HPV-2-infected sections (cutaneous warts); MAb 8C4 recognized only sections contg. HPV-16. Four MAbs (8C4, 3D1, 1D6, and 5A4) recognized a synthetic peptide corresponding to residues 269-284 of HPV-16 L1; within this region a min. antibody binding site was identified, a tripeptide 276-278. However, the complete epitope appears to extend beyond these residues and beyond HPV-16 L1 (269-284). The 5th MAb, 1C6, recognized bacterial fusion proteins contg. HPV-6b L1, -16 L1 or -18 L1 using immunoblots, yet appeared HPV-16-specific when tested on infected tissue sections. This MAb recognized 5 amino acids within a different region of HPV-16 L1 (residues 299-313).

L7 ANSWER 8 OF 10 COPYRIGHT 1993 ACS

TI Characterization of rare human papillomavirus type 11 mRNAs coding for regulatory and structural proteins, using the polymerase chain reaction

SO Virology, 172(2), 489-97

AU Rotenberg, Mitch O.; Chow, Louise T.; Broker, Thomas R.

PY 1989

AN CA111(21):188504e

AB Certain human papillomavirus (HPV) types cause warts, dysplasias, and carcinomas of the ano-genital and oral mucosa. Because of the inability to propagate HPVs in cultured cells, the paucity of viral mRNAs in human lesions, and the complexity of alternatively spliced transcripts derived from different promoters, it has not been possible to ascertain the exact structures of the majority of the mRNA species and the proteins encoded. The polymerase chain reaction was adapted to amplify cDNAs of rare, type 11 HPV mRNAs isolated from a productively infected human foreskin xenograft in an athymic mouse. The oligonucleotide primers were designed to flank each of the mRNA splice sites previously mapped by electron microscopic anal. of heteroduplexes formed between cloned HPV-11 DNA and viral mRNAs isolated from genital warts. The splice junctions were detd. by direct sequencing of the PCR-amplified cDNA products or after the cDNA was cloned into a plasmid vector. This provides the first direct evidence for the existence of rare mRNAs with the potential to encode regulatory proteins that have been hypothesized to exist for HPVs. Depending on the lengths of the upstream exons, the translation frame used and the possibility of internal reinitiation during translation, one pair of mRNAs with the same splice junction could encode the viral DNA copy no. modulating protein E1-M, the enhancer repression protein E2-C, or both. A second pair of mRNAs, also with identical splice junctions, encode the enhancer-regulating protein E2; the longer of the 2 could also encode, in its 5' exon, either or both the E6 and E7 proteins. Finally, the doubly spliced late message for the major virion capsid protein L1 also contains the entire coding region for the early E1-E4 protein in the first 2 exons, with the initiation codon for the L1 protein located precisely at the splice acceptor of the third exon. The potential of this late mRNA to encode both the E1-E4 protein and the capsid protein could contribute to the preponderance of the E4 protein in the lesion.

L7 ANSWER 9 OF 10 COPYRIGHT 1993 ACS

TI Reactivities of polyclonal and monoclonal antibodies raised to the major capsid protein of human papillomavirus type 16

SO J. Gen. Virol., 70(1), 69-77

AU Patel, Daksha; Shepherd, Philip S.; Naylor, Jennifer A.; McCance, Dennis J.

PY 1989

AN CA110(11):93221a

AB Polyclonal and monoclonal antibodies were raised against a fusion protein contg. .beta.-galactosidase and part of the major capsid protein L1 of the human papillomavirus (HPV) type 16. The polyclonal antibodies cross-reacted with the L1 protein of several HPV types including HPV-1, -2, -6 and -11 when reacted with virus-infected tissue sections, and with HPV-6 and -18 L1 fusion proteins on Western blotting. Monoclonal antibodies against the L1 fusion protein of HPV-16 reacted only with HPV-16 L1 fusion proteins on Western blots and with HPV-16-contg. biopsy sections as assessed by in situ DNA-DNA hybridization. These antibodies did not detect HPV-6 L1 protein after Western blotting or in HPV-6-infected tissue sections, although one did react with an HPV-18 fusion protein after Western blotting. The monoclonal antibodies were able to detect HPV-16 antigens in routine formaldehyde-fixed, wax-embedded sections of cervical intraepithelial neoplasm sections. HPV-16 L1 proteins were seen in one-third of biopsies that were pos. using the polyclonal cross-reacting antisera. Polyclonal antibodies to fusion proteins contg. part of the minor capsid protein L2 of

HPV-6 or -16 appeared to be more type-specific as no cross-reactivity was seen when these antibodies were reacted with HPV-1- and -2-infected tissue sections.

L7 ANSWER 10 OF 10 COPYRIGHT 1993 ACS
TI Identification of the bovine papillomavirus L1 gene
product using monoclonal antibodies
SO Virology, 165(2), 613-15
AU Cowser, Lex M.; Pilacinski, W. P.; Jenson, A. Bennett
PY 1988
AN CA109(17):146101z
AB Monoclonal antibodies (MoAbs) produced against SDS-disrupted bovine papillomavirus type 1 (BPV-1) were used to identify the product of the L1 open reading frame (ORF) of BPV-1. MoAbs were tested in ELISA with purified BPV-1 major capsid protein (MCP), fusion proteins from 2 constructions of the BPV-1 L1 ORF, and 1 construction of the L2 ORF. All MoAbs reacted with purified MCP and both L1 fusion proteins. No MoAbs were reactive with the L2 fusion protein. Polyclonal antisera raised against SDS-disrupted BPV-1 were immunoreactive with both L1 and the L2 fusion proteins. These data show that the L1 ORF of BPV-1 encodes, at least in part, the MCP of BPV-1. Further, it has been shown that the L1 encodes the papillomavirus (PV) genus-specific epitope, PV broadly cross-reactive epitope, BPV minimally cross-reactive epitope, and a BPV-1 type-specific epitope.

=> s (pv or hpv?)/ia and capsid/ia and "l1"/ia
787 PV/BI
2265 PV/AB
2661 PV/IA
(PV/BI,AB)
198 HPV?/BI
897 HPV?/AB
912 HPV?/IA
(HPV?/BI,AB)
1554 CAPSID/BI
2930 CAPSID/AB
3348 CAPSID/IA
(CAPSID/BI,AB)
1143 "L1"/BI
7564 "L1"/AB
7783 "L1"/IA
("L1"/BI,AB)
L8 36 (PV OR HPV?)/IA AND CAPSID/IA AND "L1"/IA

=> s 18 not 17
L9 26 L8 NOT L7

=> d 1-26 .mh

L9 ANSWER 1 OF 26 COPYRIGHT 1993 ACS
TI Late promoter of human papillomavirus type 8 and its regulation
SO J. Virol., 66(6), 3485-93
AU Stubenrauch, Frank; Malejczyk, Jacek; Fuchs, Pawel G.; Pfister, Herbert
PY 1992
AN CA117(5):41691d
AB Human papillomavirus type 8 (HPV8) belongs to the HPV types assocd. with skin carcinomas of patients with epidermodysplasia verruciformis (EV). Its noncoding regulatory sequences (NCR) were shown to drive the expression of the reporter gene chloramphenicol acetyltransferase (cat) in transient

assays with human epithelial cells (HT3 cells). This constitutive activity could be enhanced by coexpression of the HPV8 transactivator protein E2. The anal. of 5' deletions of the NCR showed that the EV-specific sequence motif M33 and the neighboring AP1 site are essential for the promoter activity, whereas 44 nucleotides located immediately upstream of M33 are strongly inhibitory. The same effects were obsd. in SV40 virus-immortalized fetal keratinocytes (SV61 cells) and spontaneously immortalized skin keratinocytes (HaCaT cells). By using primer extension and RNase protection analyses, 2 promoters could be identified within the HPV8 NCR. A nested set of weak signals, corresponding to start sites between positions 175 to 179, represented the previously described E6 promoter. The vast majority of transcripts was initiated at position 7535 and shown to undergo processing at an NCR-internal splice donor (positions 1 to 8). The promoter P7535 is similar to late promoters of other skin-assocd. papillomaviruses as far as localization, transcript structure, and sequence characteristics are concerned. To confirm that P7535-initiated transcripts proceed indeed to the L1 gene for the major capsid protein, viral mRNAs from an HPV8-induced lesion of a patient with EV were characterized by RNase protection and sequence anal. of polymerase chain reaction-amplified cDNAs. The NCR leader (positions 7535 to 4) appeared in 2 messages with 3 exons each. The third exon started with the second ATG codon of L1 in both cases; the short central exons from the 3' part of the early coding region were defined by a common splice acceptor site (position 3303) and different splice donor sites (positions 3443 and 3704).

L9 ANSWER 2 OF 26 COPYRIGHT 1993 ACS

TI HPV-16 viral transcripts in vulvar neoplasia: preliminary studies

SO Gynecol. Oncol., 42(3), 250-5

AU Park, J. S.; Rader, J. S.; Wu, T. C.; Laimins, L. A.; Currie, J. L.; Kurman, R. J.; Shah, K. V.

PY 1991

AN CA116(13):124779u

AB Specific human papillomavirus (HPV) types are strongly assocd. with intraepithelial neoplasia and invasive cancer of the uterine cervix. The role of HPVs in the pathogenesis of invasive carcinoma of the vulva is poorly understood. In situ hybridization for the detection of subgenomic transcripts was used in 4 vulvar specimens to elucidate the role of HPV in women. The transcripts of the E6-E7 region were more abundant than those of the L1-L2 region in vulvar neoplastic tissues. The transcripts from the early and late regions of HPV-16 continued to increase with the differentiation of the epithelial cells in both the warty and the basaloid types of vulvar precancerous lesions. This pattern persisted in invasive warty carcinomas but not in basaloid invasive carcinomas. The transcripts in basaloid carcinoma were distributed in an even and discrete pattern. L1-L2-region transcripts, as well as viral capsid protein, were detected in focal areas of well-differentiated cells of invasive warty carcinoma. The expression of HPV-16 may be regulated by the degree of cellular differentiation.

L9 ANSWER 3 OF 26 COPYRIGHT 1993 ACS

TI Detection of genital papillomavirus types by polymerase chain reaction using common primers

SO APMIS, 99(7), 667-73

AU Jenkins, Andrew; Kristiansen, Bjoern Erik; Ask, Eirik; Oskarsen, Bente; Kristiansen, Ewy; Lindqvist, Bjoern; Trope, Claes; Kjoerstad,

Kjell
 PY 1991
 AN CA116(13):121986y
 AB Eight genital human papillomavirus (HPV) types, including HPV16 and HPV18, were detected by PCR amplification of a 323-base-pair region of the genome within the L1 open reading frame (ORF). The primer sequences are: TGTAATATCCGATTWTWT and GTATCWACMACAGTAACAAA. The method will detect purified HPV16 DNA down to a concn. of as little as a single mol. in 100 .mu.L. The method is also applicable to purified DNA and crude lysates from tumor biopsies. Typing of the PCR product can be achieved with specific oligonucleotide probes.

L9 ANSWER 4 OF 26 COPYRIGHT 1993 ACS
 TI Baculovirus expression of the human papillomavirus type 16 capsid proteins: detection of L1-L2 protein complexes
 SO J. Gen. Virol., 72(12), 2981-8
 AU Xi, Shang Zhong; Banks, Lawrence M.
 PY 1991
 AN CA116(7):52585c
 AB The human papillomavirus (HPV) type 16 major capsid proteins L1 and L2 have been produced in a baculovirus expression system. Both proteins are expressed at a high level and can be readily solubilized. The L1 capsid protein migrates close to its expected Mr of 60 kDa. On the other hand L2 exhibits a much higher Mr migrating at 73 kDa, which is considerably greater than its predicted Mr of 50 kDa. The identity of both proteins has been confirmed also by Western blot anal. Both proteins are produced in drastically reduced amts. in the presence of tunicamycin. In addn. both L1 and L2 show interesting patterns of phosphorylation. L1 is phosphorylated only weakly and this appears to be quite labile, whereas L2 is very heavily phosphorylated and this, in contrast, appears to be very stable. A dual expression vector also was used for co-expressing the L1 and L2 proteins within the same baculovirus-infected cell. The results obtained from this system demonstrate the presence of protein complexes forming between the two capsid proteins. These studies indicate that at least the initial events in capsid assembly of HPVs can occur in the absence of viral DNA.

L9 ANSWER 5 OF 26 COPYRIGHT 1993 ACS
 TI Antigenic and immunogenic epitopes shared by human papillomavirus type 16 and bovine, canine, and avian papillomaviruses
 SO J. Virol., 65(12), 6862-71
 AU Dillner, Lena; Heino, Pirkko; Moreno-Lopez, Jorge; Dillner, Joakim
 PY 1991
 AN CA116(1):4843p
 AB All types of papillomaviruses (PV) share common, so-called group-specific epitopes. To identify the major group-specific epitopes, the authors immunized 26 guinea pigs or rabbits with purified bovine PV type 1 (BPV), canine PV, or avian PV from the common chaffinch. The resulting hyperimmune sera, as well as a com. available rabbit antiserum to BPV and seven monoclonal antibodies to BPV, were tested in an ELISA with a set of 66 overlapping 20-amino-acid peptides representing the complete sequence of the major capsid proteins (L1 and L2) of human PV type 16 (HPV 16). Sera from the same animals before immunization were used as controls. The minimal reactive epitopes within each peptide were further characterized by testing of truncated peptides. The

cross-reactive epitopes were clustered in the regions of L1, an internal region (at positions 171 to 235), which contained three epitopes, and the more reactive region at the carboxy terminus (at positions 411 to 475), which contained six epitopes. The most reactive of the HPV 16 broadly cross-reactive epitopes was a carboxy-terminal epitope which had the sequence DTYRF and which reacted with nine of the antisera to BPV, canine PV, or avian PV, with the com. available rabbit antiserum to BPV, and also with a mouse monoclonal antibody to BPV. Antipeptide antisera to all of the HPV 16 L1 peptides and to the most antigenically reactive of their truncated analogs were made in guinea pigs. Antipeptide antisera reactive with BPV were obtained for three of the cross-reactive epitopes, and one of these antisera allowed highly sensitive detection of group-specific PV antigen by immunoperoxidase staining.

- L9 ANSWER 6 OF 26 COPYRIGHT 1993 ACS
 TI Expression of vaccinia recombinant HPV 16 L1 and
L2 ORF proteins in epithelial cells is sufficient for assembly of
HPV virion-like particles
 SO Virology, 185(1), 251-7
 AU Zhou, Jian; Sun, Xiao Yi; Stenzel, Deborah J.; Frazer, Ian H.
 PY 1991
 AN CA115(23):251998u
 AB A recombinant vaccinia virus termed pLC201VV was designed to
 coexpress the L1 and L2 late genes of human papillomavirus
 type 16 (HPV16). Synthesis of the L1 and L2
 proteins occurred in cells infected with pLC201VV, and 40-nm
 virus-like particles with a d. of 1.31 g/mL were produced in the
 nuclei of cells synthesizing both L1 and L2, but not in
 cells synthesizing either protein alone. Virus-like particles were
 partially purified from infected cells by sucrose gradient
 sedimentation and shown to consist of capsomeres similar to
HPV and contain glycosylated L1 viral
capsid protein. The prodn. of HPV-like particles
 using recombinant vaccinia virus should be useful for biochem.
 studies and could provide a safe source of material for the
 development of a vaccine.
- L9 ANSWER 7 OF 26 COPYRIGHT 1993 ACS
 TI Type-specific and cross-reactive epitopes in human papillomavirus
 type 16 capsid proteins
 SO Virology, 184(1), 460-4
 AU Beiss, Barbara K.; Heimer, Edgar; Felix, Arthur; Burk, Robert D.;
 Ritter, Diane B.; Mallon, Robert G.; Kadish, Anna S.
 PY 1991
 AN CA115(17):181030w
 AB Rabbit polyclonal and mouse monoclonal antisera were raised to C
 terminal peptides from the genital human papillomavirus (HPV
) 16 L1 and L2 open reading frames (ORFs). Anti-L1
 and -L2 peptide sera recognized HPV 16 L1 and L2
 fusion proteins in Western blots and by immunopptn. In Western blot
 anal. of L1 proteins from different HPV types,
 antisera to the L1 peptide reacted only with HPV
 16, thus identifying an HPV 16 type-specific linear
 epitope. Anti-L2 peptide sera reacted with L2 fusion proteins from
HPVs 6 and 16, but not from BPV, thus identifying a
 partially cross-reactive epitope in the HPV 16 L2.
 Computer anal. of C terminal amino acid sequences of the L1
 and L2 ORFs of multiple HPV types supported the Western
 blot findings. Despite the HPV 16 type specificity found
 in Western blots, anti-L1 peptide sera identified nuclear

antigen by immunocytochem. in cervical biopsies infected with HPV 16, as well as other genital HPV types.

Anti-L2 peptide sera failed to recognize antigen in infected tissue.

L9 ANSWER 8 OF 26 COPYRIGHT 1993 ACS

TI Expression of human papillomavirus proteins in yeast *Saccharomyces cerevisiae*

SO Virology, 182(2), 513-21

AU Carter, Joseph J.; Yaegashi, Nobuo; Jenison, Steven A.; Galloway, Denise A.

PY 1991

AN CA115(1):2410a

AB The L1 and L2 proteins of human papillomavirus (HPV) types 1, 6, and 16 and the E6 and E7 proteins of HPV 16 were expressed in *S. cerevisiae*. The yeast expressed proteins were readily detected by immune blotting and were generally intact. The HPV 1 L1 and L2 proteins expressed in yeast were indistinguishable from the major and minor capsid proteins purified from HPV 1 virions as judged by gel electrophoresis and immunoblotting. The HPV 6 and HPV 16 L2 proteins and HPV 16 E7 proteins were secreted from yeast by fusion to the yeast pre-pro- α -factor leader sequence. Following secretion of the HPV 16 E7 protein a rapid method of purification was developed. The yeast expressed proteins were used as antigen targets to study the human immune response in Western blot assay, ELISA, and immune precipitation. One human serum reacted with intact, but not denatured HPV 16 L2 proteins, suggesting that the yeast expressed proteins will be useful to detect antibodies reactive with conformational epitopes.

L9 ANSWER 9 OF 26 COPYRIGHT 1993 ACS

TI Nucleotide sequence of human papillomavirus (HPV) type 41: an unusual HPV type without a typical E2 binding site consensus sequence

SO Virus Res., 18(2-3), 179-89

AU Hirt, Lorenz; Hirsch-Behnam, Anja; De Villiers, Ethel Michele

PY 1991

AN CA115(1):2028g

AB The complete nucleotide sequence of human papillomavirus type 41 (HPV-41) has been determined. HPV-41 was originally isolated from a facial wart, but its DNA has subsequently been detected in some skin carcinomas and premalignant keratoses (Grimmel, M., et al., 1988, and E.-M. de Villiers, M. Grimmel, and C. Neumann, unpublished results). Analysis of the cloned HPV -41 nucleic acid reveals that its genome organization is similar to that of other papillomavirus types. Yet, the analysis indicates at the same time that this virus is most distantly related to all other types of human-pathogenic papillomaviruses sequenced thus far and appears to identify HPV-41 as the first member of a new subgroup of HPV. The overall nucleotide homology to other sequenced HPV types is below 50%. The closest other HPV type is represented by HPV-18, sharing 49% identical nucleotides. The typical E2-binding sequence ACCN6GGT, found in all papillomaviruses analyzed to date, does not occur in the upper regulatory region of the HPV-41 genome. Modified E2-binding sequences, as described for BPV 1 (Li, R., et al., 1989), are located in the domain proximal to the E6 ORF. These are ACCN6GTT, AACN6GGT and the 2 perfect palindromic sequences AACGAATTCGTT.

L9 ANSWER 10 OF 26 COPYRIGHT 1993 ACS

TI The open reading frame L2 of cottontail rabbit papillomavirus

contains antibody inducing neutralizing epitopes
SO Virology, 181(2), 572-9
AU Christensen, Neil D.; Kreider, John W.; Kan, Nancy C.; DiAngelo,
Susan L.
PY 1991
AN CA114(17):162137z
AB Polyclonal antisera were generated against bacterially derived
fusion proteins of the open reading frames (ORFs) of the
capsid proteins of cottontail rabbit papillomavirus (CRPV).
The carboxy-terminal two-thirds of CRPV L1 and the
carboxy-terminal half of CRPV L2 were cloned into a bacterial
expression vector and induced proteins were used as antigen and
immunogen. The polyclonal antisera were tested in a series of
immunol. assays, including ELISA, Western blot, and neutralization
of CRPV. ELISA demonstrated that the polyclonal antisera raised
against expressed L1 proteins reacted strongly to
disrupted CRPV virion antigen and weakly both to intact CRPV virion
and disrupted BPV-1 (bovine papillomavirus 1) virion. Anti-CRPV L2
antisera reacted strongly only to intact and disrupted CRPV virion
antigen. Viral capsid proteins of CRPV were detected in
Western blots of human HPV-11, BPV-1, and CRPV virus
particles by these polyclonal antisera. The anti-L1 sera
recognized the major capsid protein (60 kDa) and the
anti-L2 sera identified a 76 kDa viral protein of CRPV. Only the
antisera generated against expressed L2 neutralized CRPV. The
neutralizing titer of the anti-L2 sera, however, was several orders
of magnitude lower than the titer of a neutralizing polyclonal
antiserum that was generated by immunizations with intact CRPV
virions.

L9 ANSWER 11 OF 26 COPYRIGHT 1993 ACS
TI Definition of murine T helper cell determinants in the major
capsid protein of human papillomavirus type 16
SO J. Gen. Virol., 71(11), 2691-8
AU Davies, D. Huw; Hill, C. Mark; Rothbard, Jonathan B.; Chain,
Benjamin M.
PY 1990
AN CA114(5):40580t
AB Three murine major histocompatibility complex (MHC) class
II-restricted T cell determinants were identified in the major
capsid protein L1 of human papillomavirus (
HPV) type 16. Peptides derived from HPV-16
L1, which contain putative T cell epitopes located by a
predictive algorithm, were synthesized and tested for
lymphoproliferative activity by direct immunization, followed by in
vitro assay of responses to peptides or recombinant HPV-16
L1. The MHC restriction of the stimulatory peptides was
detd. using blocking monoclonal antibodies against class II mols.
The responses, which were specific for the priming peptides alone,
cross-reacted with recombinant L1 but not with analogous
peptides derived from other HPV types.

L9 ANSWER 12 OF 26 COPYRIGHT 1993 ACS
TI Identification of seroreactive regions of the human papillomavirus
type 16 proteins E4, E6, E7 and L1
SO J. Gen. Virol., 71(11), 2709-17
AU Mueller, Martin; Gausepohl, Heinrich; De Martynoff, Guy; Frank,
Rainer; Brasseur, Robert; Gissmann, Lutz
PY 1990
AN CA113(25):229160b
AB Small fragments of the DNA of human papillomavirus type 16 (
HPV-16) were randomly cloned into the bacteriophage fd which

expresses the resulting peptides as part of its capsid. Antisera raised against different HPV-16 fusion proteins were used for screening of the phage clones and the reacting peptides were detd. by sequencing the inserted HPV-16 DNA fragments of the pos. recombinants. Seroreactive regions of the proteins derived from the E4, E6, E7 (two regions) and L1 (three regions) open reading frames could be found by this approach. Of these seven regions, four were defined by at least two overlapping inserts, thus limiting the domains to between 10 and 15 amino acids. In the case of the E4 open reading frame, the same region identified by immunoscreening was also found when synthetic overlapping octapeptides were tested by ELISA with the anti-E4 antiserum. Using an approach to predict receptor-like regions within the resp. proteins, five of the seven regions were also identified. From the data on these regions, synthetic peptides were produced and used for the detection of antibodies against HPV-16 proteins in human sera by ELISA.

L9 ANSWER 13 OF 26 COPYRIGHT 1993 ACS

TI Immunochemical method for detection of human papillomavirus antibodies, peptides useful in the method, and use of the method for diagnosis, especially of cervical carcinoma

SO PCT Int. Appl., 57 pp.

AU Dillner, Joakim; Dillner, Lena

PI WO 9004790 A1 3 May 1990

AI WO 89-SE612 30 Oct 1989

PY 1990

AN CA113(13):113688a

AB A method is provided for detection of human papillomavirus (HPV) for diagnosis, esp. for diagnosis of carcinoma or pre-stages thereof, or the risk of development of carcinoma. The method relies on detecting the presence of IgA, IgG, and IgM antibodies against papillomavirus virions in a body fluid, esp. a cervical secretion. The virions include individual virion proteins or peptides thereof. Thus, 66 peptides (20 amino acid residues each) with a 5 residue overlap to each other were synthesized according to the deduced amino acid sequences of the L1 and L2 open reading frames (encoding viral capsid proteins) for HPV16. The peptides were used in an ELISA testing sera from HPV16-carrying cervical neoplasia patients for reactivity with either IgA, IgG, or IgM. Reactivity for individual serum samples using individual peptides is shown. The 7 most immunoreactive peptides were also tested for IgA, IgG, and IgM reactivity in 60 control serum samples, derived from healthy donors or patients with irrelevant tumors. Most of these peptides showed significant immunoreactivity only with <10% of the control sera.

L9 ANSWER 14 OF 26 COPYRIGHT 1993 ACS

TI An antigen chimera of poliovirus induces antibodies against human papillomavirus type 16

SO J. Virol., 64(3), 1201-6

AU Jenkins, Owen; Cason, John; Burke, Karen L.; Lunney, Declan; Gillen, Alison; Patel, Daksha; McCance, Dennis J.; Almond, Jeffrey W.

PY 1990

AN CA113(3):21849a

AB It has been established that the surface of poliovirus type 1 can be extensively modified to incorporate antigenic domains from other poliovirus serotypes and from unrelated viruses. The fact that the modified (chimeric) viruses exhibit dual antigenicity and immunogenicity led to exploring the possibility of using the Sabin vaccine strain of poliovirus type 1 as a vector for the presentation of antigenic domains from human papillomavirus type 16 (HPV

-16), a virus a●cd. with the development ● cervical carcinoma. This report describes the construction and characterization of a chimeric poliovirus contg. a 16-residue sequence derived from the major capsid protein (L1) of HPV-16.

This virus chimera stimulated the prodn. in rabbits of antibodies which recognized the HPV-16-derived peptide and an L1 fusion protein synthesized in Escherichia coli and detected HPV-16 in human biopsy material by immunoperoxidase staining. The possibility that poliovirus-HPV chimeras could be used as vaccines against HPV-16 is discussed.

L9 ANSWER 15 OF 26 COPYRIGHT 1993 ACS

TI Mapping of linear epitopes of human papillomavirus type 16: the L1 and L2 open reading frames

SO Int. J. Cancer, 45(3), 529-35

AU Dillner, Joakim; Dillner, Lena; Utter, Goeran; Eklund, Carina; Rotola, Antonella; Costa, Silvano; DiLuca, Dario

PY 1990

AN CA113(1):1232r

AB Certain types of human papillomavirus (HPV), notably HPV type 16, are assocd. with flat or inverted proliferative lesions of the cervix uteri that can progress to malignancy. As a first step towards the serol. study of the epidemiol. of HPV, the entire amino acid sequences of the 2 major viral capsid proteins of HPV type 16, L1 and L2 were synthesized, as a set of 66 synthetic 20-residue peptides with an overlap of 5 amino acids. The peptides were tested for reactivity with IgA, IgG and IgM antibodies in the sera of 30 patients with HPV-16-carrying cervical neoplasms. Both IgG and IgM antibody responses were detected, but most of the reactivity found was of the IgA class. The most immunoreactive peptides were further analyzed for reactivity with sera from 22 patients with parotid gland tumors and with sera from 38 healthy individuals. The L2-encoded protein contained only one major linear epitope, which was not specific for HPV-16-carrying neoplasms. In contrast, the L1-encoded protein contained several epitopes that were regularly immunoreactive with antibodies present in the sera of patients with HPV-16-carrying cervical neoplasms, but only rarely so in the sera of patients with other tumors or of healthy individuals.

L9 ANSWER 16 OF 26 COPYRIGHT 1993 ACS

TI Human T cell responses to human papillomavirus type 16 L1 and E6 synthetic peptides: identification of T cell determinants, HLA-DR restriction and virus type specificity

SO J. Gen. Virol., 71(2), 423-31

AU Strang, George; Hickling, Julian K.; McIndoe, G. Angus J.; Howland, Kevin; Wilkinson, David; Ikeda, Hitoshi; Rothbard, Jonathan B.

PY 1990

AN CA112(23):214980z

AB Four T cell determinants in the major capsid protein of human papillomavirus (HPV) type 16 L1 and one in the E6 protein assocd. with cellular transformation were defined using synthetic peptides to stimulate peripheral blood mononuclear cells from asymptomatic individuals. HLA-DR restriction was defined using murine L cells transfected with HLA-DR genes to present antigen. Responses to two of the five determinants by T cell lines and clones were shown to be specific for HPV-16 based on the lack of cross-recognition of the corresponding sequences of other known papillomavirus sequences (types 1a, 5, 6b, 8, 11, 18, and 33). The T cells raised against two of the other peptides

cross-reacted with corresponding peptides from other strains to varying extents, depending on their structural homol. The implications of these results regarding the prevalence of HPV-16 infection in the population and the possible diagnostic role of these responses in papillomavirus infection is discussed.

L9 ANSWER 17 OF 26 COPYRIGHT 1993 ACS

TI Immunological cross-reactivity to laboratory-produced HPV -11 virions of polysera raised against bacterially derived fusion proteins and synthetic peptides of HPV-6b and HPV -16 capsid proteins

SO Virology, 175(1), 1-9

AU Christensen, Neil D.; Kreider, John W.; Cladel, Nancy M.; Galloway, Denise A.

PY 1990

AN CA112(23):214795t

AB Polysera raised in rabbits to bacterially derived fusion proteins and synthetic peptides of the L1 and L2 ORFs of human papillomaviruses (HPV)-6b and -16 were tested for cross-reactivity to lab.-produced infectious HPV-11 virions. The polysera were analyzed in a series of 5 different immunol. assays including immunoperoxidase staining of the koilocytotic nuclei in sections of formalin-fixed, paraffin-embedded as well as fresh frozen sections of HPV-11 expt1. condylomas generated in the athymic nude mouse xenograft system, ELISA, Western blots, and neutralization of infectious HPV -11 virions. ELISA and Western blot assays were used to det. whether the polysera identified external or internal epitopes on HPV -11 virions, and whether there was cross-reactivity to bovine papillomavirus-1 or lab.-produced cottontail rabbit papillomavirus virions. Seven of a total of 12 sera were pos. for reactivity to HPV-11 in one or more assays, but none of the reactivity was directed to external epitopes on the intact virions as detd. by ELISA. None of the L1 products generated group-specific antigen (GSA) antisera including a synthetic peptide spanning the GSA site. The combination of assays clearly demonstrated that apparent false pos. and false neg. reactivities of different antisera were obtained for each assay system tested. Thus, no single assay could be used reliably to det. the true antiviral reactivity of a given polysera.

L9 ANSWER 18 OF 26 COPYRIGHT 1993 ACS

TI Humoral assays of human sera to disrupted and nondisrupted epitopes of human papillomavirus type 1

SO Virology, 174(2), 388-98

AU Steele, Jane C.; Gallimore, Phillip H.

PY 1990

AN CA112(15):136991n

AB The use of different assay systems and the disparity in results obtained has meant that there is little understanding about the role played by the humoral response during human papillomavirus (HPV) infection. Human antibody responses have so far appeared to be largely directed against the major capsid protein, L1. This protein possesses both type-specific and type-common antigenic determinants but it is not known which of these is important in vivo during the natural course of infection. In this study humoral responses of individuals to purified HPV 1 virions were tested in 3 types of antibody assay. Western blot anal. detected antibodies in only 8 of 83 serum samples, whereas an ELISA and immunopptn. assay using nondisrupted HPV 1 virions showed pos. antibody reactivities for 71 and

64 individuals, esp. It is suggested that the humoral response to L1 is mainly directed against native conformational epitopes present on the whole HPV 1 particle and that type-common epitopes are not largely involved. This was further demonstrated by the fact that when samples were tested in the same ELISA system using disrupted HPV 1 virions as the antigen instead of whole virus particles, the no. of pos. sera was reduced to 9 out of 83. Thus, humoral assays using antigenic material pertaining to disrupted HPV epitopes are of limited use, at least in the case of HPV type 1. There were no obvious correlations between the antibody assay results and clin. histories of wart infection except that a lower no. of pos. serum reactivities were found among the group of individuals claiming to have no past history of HPV infection.

L9 ANSWER 19 OF 26 COPYRIGHT 1993 ACS

TI Sensitive detection of nucleic acids and protein of human papillomavirus type 6 in respiratory and genital tract papillomata

SO J. Virol. Methods, 25(1), 31-47

AU Wu, Tzyy Choou; Mounts, Phoebe

PY 1989

AN CA111(25):228424n

AB A sensitive method was developed to detect and localize HPV-6 viral DNA, mRNA, and protein in biopsy specimens of genital and respiratory tract lesions by using in situ hybridization and immunoperoxidase assays on sections of plastic embedded tissue. This modified in situ hybridization technique, using ultrathin sections and strand-specific 3H-labeled riboprobes, offers the advantages of superior morphol. preservation and detection of viral genomes at low copy no. with good resolu. This modified immunocytochem. provides better sensitivity when compared to previous methods using paraffin-embedded materials. In respiratory tract lesions, immunoperoxidase assay detected only a few capsid antigen-pos. cells, while in the genital tract lesions, there were more capsid antigen-pos. cells. Southern transfer analyses and in situ hybridizations demonstrated the presence of more viral nucleic acids in genital tract papillomata than respiratory tract papillomata. Epithelial cells throughout the papillomata were infected by HPV-6 as evidenced by pos. hybridization, with more viral DNA present in superficial cells. Apparently, genital tract epithelium is more permissive for HPV-6 replication than respiratory tract epithelium. Using stand-specific probes synthesized from subgenomic fragments of the HPV-6 genome in conjunction with nuclease digestions, it was possible to demonstrate that HPV-6 transcripts specific to open reading frames (ORFs) E6, E7, E1, L1, and L2 occur in maturing superficial cells. In contrast, transcripts specific to ORFs E1, E2, E4, E5a, and E5b could be detected throughout the whole of the epithelium with more signals noted at the basal cell areas. In addn., the distribution of HPV-6 nucleic acids and protein in a carcinoma in situ of the larynx was analyzed. In comparison to benign respiratory tract papillomata, more viral DNA was found in the malignant lesion, but the pattern and distribution of transcription and capsid antigen was similar.

L9 ANSWER 20 OF 26 COPYRIGHT 1993 ACS

TI Differentiation-linked human papillomavirus types 6 and 11 transcription in genital condylomata revealed by in situ hybridization with message-specific RNA probes

SO Virology, 172(1), 331-40

AU Stoler, Mark H.; Wolinsky, Steven M.; Whitbeck, April; Broker, Thomas R.; Chow, Louise T.

PY 1989
AN CA111(19):168253u
AB Human papillomaviruses (HPVs) infect specific human epithelial tissues. Because viral propagation in cultured cells has not been achieved, studies of HPV genetic activities have been difficult and rely largely on analyses of patient specimens by conventional biochem. methods. HPV type 6 and type 11 infections often result in genital warts (condylomata acuminata). Structural mapping of RNAs from such warts reveals that they use alternative promoters, splice sites, and polyadenylation sites to produce complex families of overlapping mRNAs that span multiple open reading frames. Based on the mRNA structures, message-specific subgenomic clones of HPV-6 and HPV-11 were developed in pGEM vectors. Tritium-labeled, single-stranded RNA probes were synthesized in vitro and applied to serial thin sections of patient specimens for in situ hybridization. The data show the qual. and quant. transcription patterns of different viral messages in relationship to one another, to viral DNA replication, and to cellular differentiation. The viral E region is transcribed before the onset of vegetative DNA replication and continues to be expressed in increasing amts. in the maturing epithelium. Even in mature epithelia, E region messengers are far more abundant than L region mRNAs. The L region messages encoding capsid proteins are truly late in that they appear concomitant with or after the onset of vegetative viral DNA replication and are only present in the superficial strata of the epithelium, which contain the oldest and most differentiated keratinocytes. Abundant intron material derived from processing E region transcripts accumulates in the nuclei. Strictly nuclear signals from the L region transcripts in the midepithelium suggest that regulation of their expression is at the level of transcription elongation.

L9 ANSWER 21 OF 26 COPYRIGHT 1993 ACS
TI Expression in Escherichia coli of seven DNA fragments comprising the complete L1 and L2 open reading frames of human papillomavirus type 6b and localization of the 'common antigen' region
SO J. Gen. Virol., 70(3), 543-55
AU Strike, David G.; Bonneze, William; Rose, Robert C.; Reichman, Richard C.
PY 1989
AN CA110(19):167069f
AB Mol. cloning was used to express human papillomavirus type 6b (HPV-6b) antigens in E. coli. Seven genomic DNA fragments of HPV-6b which together comprise the complete L1 and L2 open reading frames, known to code for capsid proteins, were cloned and expressed in E. coli as both .beta.-galactosidase and TrpE fusion proteins. Western blots of HPV-6b .beta.-galactosidase fusion proteins using genus-specific antisera produced by immunization of rabbits with disrupted bovine papillomavirus type 1 (BPV-1) showed that polypeptides encoded by two DNA fragments from the mid portion of L1 of HPV-6b were cross-reactive. Only one of these two polypeptides reacted with antisera raised against disrupted HPV-1, directly demonstrating that this polypeptide contains the papillomavirus common antigen. The cross-reactive region was confirmed by reversing antigen and antibody. Polyclonal antisera were raised against the seven HPV-6b .beta.-galactosidase fusion proteins and tested against BPV-1 virion proteins on Western blots. Only antiserum against the mid portion of L1 of HPV-6b reacted with BPV-1 major capsid protein. HPV-6b fusion

proteins were also used to test human sera for antibodies reactive in Western blots. Serum samples from 38 patients with documented HPV-6 infections and from 22 presumably uninfected controls were tested. Antibodies were not detected in any of the sera to any of the seven fusion proteins. HPV-6b .beta.-galactosidase fusion proteins are antigenic and can be used on Western blots to localize immunol. reactive subregions of proteins by reacting protein fragments with antisera from immunized animals. However, alternative methods will be required to detect anti-HPV antibodies in human sera.

L9 ANSWER 22 OF 26 COPYRIGHT 1993 ACS

TI Detection of human papillomavirus capsid antigens in various squamous epithelial lesions using antibodies directed against the L1 and L2 open reading frames

SO Virology, 164(2), 467-77

AU Firzlaff, Juliane M.; Kiviat, Nancy B.; Beckmann, Anna Marie; Jenison, Steven A.; Galloway, Denise A.

PY 1988

AN CA109(11):90524v

AB HPV6 and HPV16 infect the squamous epithelium of the genital tract and are thought to be involved in the pathogenesis of benign and malignant lesions. HPV6 is primarily found in benign condylomas whereas HPV16 is present in dysplasias and in invasive squamous cell carcinomas. To examine the expression of the major and minor capsid proteins in these lesions, polyclonal antisera directed against bacterially derived fusion proteins harboring different restriction fragments of the L1 and L2 ORFs of HPV6b and HPV16 were generated. L1 ORF-specific antisera were not type-specific and detected the major capsid antigen in lesions infected with related HPV types. Anti-L2 ORF antisera could distinguish among HPV1, HPV6, and HPV16 when the fusion protein used as the immunogen did not harbor the amino-terminus of the L2 ORF. The anti-L1 ORF antisera were employed to detect the major capsid protein in various lesions by immunohistochem. staining. Lesions harboring HPV16 were pos. in a high percentage of cervical intraepithelial neoplasia I-II (87%), and less frequently in carcinomas in situ (29%) or invasive carcinomas (17%). In all cases capsid antigen expression was restricted to cells showing some differentiation at the surface or periphery of the lesion.

L9 ANSWER 23 OF 26 COPYRIGHT 1993 ACS

TI Expression of human papillomavirus type 6 and type 16 capsid proteins in bacteria and their antigenic characterization

SO J. Gen. Virol., 68(12), 3081-9

AU Banks, L.; Matlashewski, G.; Pim, D.; Churcher, M.; Roberts, C.; Crawford, L.

PY 1987

AN CA108(13):109354t

AB The L1 and L2 ***capsid*** proteins encoded by human papillomavirus types 6 and 16 (HPV-6 and HPV-16) were synthesized in bacteria. Antisera were raised against the HPV-6 L1- and L2-.beta.-galactosidase fusion proteins and against an HPV-16 L1 C-terminal peptide which was 14 amino acids long. The HPV-16 L1 peptide antibodies were highly reactive with the HPV-16 L1-.beta.-galactosidase fusion protein but not against the equiv. HPV-6 L1 -.beta.-galactosidase fusion protein. The effectiveness of these antibodies was compared with com. available antiovine

papillomavirus type 1 (BPV-1) antibodies and the results demonstrated that the anti-BPV-1 antibodies reacted well against HPV-6 L1-.beta.-galactosidase but not against HPV-16 L1-.beta.-galactosidase. In addn., the L2 portion of the HPV-6 L2-.beta.-galactosidase fusion protein appeared particularly immunogenic, since antibodies raised against this fusion protein were predominantly reactive with the L2 moiety. The HPV-16 L1 peptide antibodies described here will be preferred reagents for the specific detection of HPV-16 capsid antigens, which may be particularly important in early diagnosis of HPV-16 infection.

L9 ANSWER 24 OF 26 COPYRIGHT 1993 ACS

TI Identification of the human papillomavirus type 6b L1 open reading frame protein in condylomas and corresponding antibodies in human sera

SO J. Virol., 61(9), 2684-90

AU Li, Chou Chi; Shah, Keerti V.; Seth, Arun; Gilden, Raymond V.

PY 1987

AN CA107(15):130590f

AB Genital warts (condylomata acuminata) are among the most frequent sexually transmitted infections. Human papillomavirus type 6 (

HPV-6), which is etiolo. related to a majority of these

lesions, has not been propagated in tissue culture. Two forms of

HPV-6 viral antigens were generated: a chem. synthesized

oligopeptide (referred to as the C-terminal synthetic peptide)

corresponding to residues 482 to 495 of the 500-amino-acid-long

L1 open reading frame (ORF), and a bacterially expressed

54-kilodalton (kDa) fusion protein contg. the N-terminal 13 amino

acids encoded by the .lambda. bacteriophage cII gene followed by one vector-insert junctional residue and 462 amino acids of the

L1 ORF sequence (residues 39 to 500). The cII-L1

fusion protein was specifically recognized by an antipeptide serum directed against the N-terminal 13 amino acids derived from the cII gene, an antiserum raised against the C-terminal synthetic peptide, and a genus-specific serum prepd. by immunization with disrupted viral capsids. The 54-kDa fusion protein was purified, and the sequence of its first 36 amino acids was detd. and found to be as predicted by the DNA sequence. Both the genus-specific anticapsid serum and the antiserum raised against the fusion protein identified authentic L1 ORF proteins in HPV-1-induced (58

kDa) and HPV-6/11-induced (56 kDa) papillomas. The

synthetic peptide antiserum recognized the 56- to 58-kDa protein in

HPV-6-induced warts, but not in HPV-1- or

HPV-11-infected specimens. Using the fusion protein as

antigen in immunoassays, the corresponding antibodies were detected in human sera.

L9 ANSWER 25 OF 26 COPYRIGHT 1993 ACS

TI Expression of human papillomavirus types 6b and 16 L1 open reading frames in Escherichia coli: detection of a 56,000-dalton polypeptide containing genus-specific (common) antigens

SO J. Virol., 61(8), 2389-94

AU Tomita, Yoshimi; Shirasawa, Hiroshi; Simizu, Bunsiti

PY 1987

AN CA107(11):91237z

AB The human papillomavirus (HPV) genome contains two large

open reading frames (ORFs), designated L1 and L2. To

characterize the antigenic properties of the L1

ORF-encoded proteins, the L1 ORFs of HPV6b and

HPV16 were cloned in plasmids, and these were expressed in

E. coli. First, the HPV6b DNA, representing 5.2% of the L1 ORF, was cloned in pUC19 and expressed in E. coli JM83 and RB791 as a 160,000-mol.-wt. (160K) fusion protein with E. coli .beta.-galactosidase (6bL1/.beta.-gal). Second, the HPV16 DNA, representing 89.8% of the L1 ORF, was cloned in pKK233-2 and expressed as a 56K protein (16L1) in strain RB791. Both the 6bL1/.beta.-gal and 16L1 proteins cross-reacted with anti-bovine papillomavirus type 1 (BPV1) antibody raised against disrupted BPV1 particles. An antibody raised against the 6bL1/.beta.-gal fusion protein reacted with the 16L1 protein and also with native papillomavirus antigens in human genital condyloma and bovine fibropapilloma tissues, as detd. by biotin-streptavidin staining. Furthermore, the anti-6bL1/.beta.-gal antibody recognized a 54K protein which seemed to be a major capsid protein of BPV1 and also a 56K protein of biopsies harboring HPV6 or HPV11. It was concluded that the papillomavirus L1 gene product contains genus-specific (common) antigens and that the HPV6 and HPV11 L1 genes specify the 56K capsid protein.

L9 ANSWER 26 OF 26 COPYRIGHT 1993 ACS
TI Expression of the human papillomavirus type 6b L2 open reading frame in Escherichia coli: L2-.beta.-galactosidase fusion proteins and their antigenic properties
SO Virology, 158(1), 8-14
AU Tomita, Yoshimi; Shirasawa, Hiroshi; Sekine, Hiromasa; Simizu, Bunsiti
PY 1987
AN CA106(25):208815j
AB The human papillomavirus (HPV) type 6b genome contains 2 large open reading frames (ORFs), designated L1 and L2, in a putative late region. These ORFs are expected to code for viral structural proteins. To exam. antigenic properties of a L2 gene product, two plasmids which contain N-terminal (L2-N) and internal (L2-I) regions of the HPV6b L2 ORF were constructed and then each region was expressed in E. coli as a fusion protein with E. coli .beta.-galactosidase (.beta.-Gal). Both L2-N/.beta.-Gal and L2-I/.beta.-Gal fusion proteins reacted with anti-.beta.-Gal antibody, but did not react with the antibody prep. against bovine papillomavirus type 1 (BPV1), in contrast with a high reactivity of HPV6b L1-.beta.-Gal fusion protein with the anti-BPV1 antibody. Antibody raised against the L2-I/.beta.-Gal protein in a rabbit reacted with viral antigens in the nuclei of cells in superficial epithelium of the condyloma acuminatum tissue, but did not react with the antigens in the bovine papilloma tissue. This antibody recognized a protein from condyloma acuminata which migrates to the position of mol. wt. 70K-76K on an electrophoresed SDS-polyacrylamide gel. These results suggested that the L2 ORF of HPV6b codes for a capsid protein which is less cross-reactive than the L1 antigen with anti-BPV1 antibody.

=> fil .biotech

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.. => s (papillomavir? or hpv or pv) and capsid and "l1"

FILE 'BIOSIS'

4551 PAPILLOMAVIR?
2540 HPV
3328 PV
4701 CAPSID
3365 "L1"

L10 48 (PAPILLOMAVIR? OR HPV OR PV) AND CAPSID AND "L1"

FILE 'MEDLINE'

4573 PAPILLOMAVIR?
2682 HPV
1742 PV
5866 CAPSID
3351 "L1"

L11 56 (PAPILLOMAVIR? OR HPV OR PV) AND CAPSID AND "L1"

FILE 'EMBASE'

2861 PAPILLOMAVIR?
2258 HPV
1513 PV
2445 CAPSID
2121 "L1"

L12 45 (PAPILLOMAVIR? OR HPV OR PV) AND CAPSID AND "L1"

TOTAL FOR ALL FILES

L13 149 (PAPILLOMAVIR? OR HPV OR PV) AND CAPSID AND "L1"

=> s l13 and (schlegel c? or jensen a?)/au

FILE 'BIOSIS'

11 SCHLEGEL C?/AU
639 JENSEN A?/AU

L14 0 L10 AND (SCHLEGEL C? OR JENSEN A?)/AU

FILE 'MEDLINE'

16 SCHLEGEL C?/AU
301 JENSEN A?/AU

L15 0 L11 AND (SCHLEGEL C? OR JENSEN A?)/AU

FILE 'EMBASE'

4 SCHLEGEL C?/AU
151 JENSEN A?/AU

L16 0 L12 AND (SCHLEGEL C? OR JENSEN A?)/AU

TOTAL FOR ALL FILES

L17 0 L13 AND (SCHLEGEL C? OR JENSEN A?)/AU

=> s (schlegel c? and jensen a?)/au

FILE 'BIOSIS'

11 SCHLEGEL C?/AU
639 JENSEN A?/AU

L18 0 (SCHLEGEL C? AND JENSEN A?)/AU

FILE 'MEDLINE'

16 SCHLEGEL C?/AU
301 JENSEN A?/AU

L19 0 (SCHLEGEL C? AND JENSEN A?)/AU

FILE 'EMBASE'

4 SCHLEGEL C?/AU
151 JENSEN A?/AU

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L20      0 (SCHLEGEL C? AND JENSEN A?)/AU
TOTAL FOR ALL FILES
L21      0 (SCHLEGEL C? AND JENSEN A?)/AU

=> s (papillomavir? or hpv or pv) and capsid and "l1 protein"
FILE 'BIOSIS'
      4551 PAPILOMAVIR?
      2540 HPV
      3328 PV
      4701 CAPSID
      3365 "L1"
      559680 "PROTEIN"
      51 "L1 PROTEIN"
      ("L1"(W)"PROTEIN")
L22      16 (PAPILOMAVIR? OR HPV OR PV) AND CAPSID AND "L1 PROTEIN"

FILE 'MEDLINE'
      4573 PAPILOMAVIR?
      2682 HPV
      1742 PV
      5866 CAPSID
      3351 "L1"
      414944 "PROTEIN"
      46 "L1 PROTEIN"
      ("L1"(W)"PROTEIN")
L23      13 (PAPILOMAVIR? OR HPV OR PV) AND CAPSID AND "L1 PROTEIN"

FILE 'EMBASE'
      2861 PAPILOMAVIR?
      2258 HPV
      1513 PV
      2445 CAPSID
      2121 "L1"
      280799 "PROTEIN"
      36 "L1 PROTEIN"
      ("L1"(W)"PROTEIN")
L24      11 (PAPILOMAVIR? OR HPV OR PV) AND CAPSID AND "L1 PROTEIN"

TOTAL FOR ALL FILES
L25      40 (PAPILOMAVIR? OR HPV OR PV) AND CAPSID AND "L1 PROTEIN"

=> dup rem l25
PROCESSING COMPLETED FOR L25
L26      19 DUP REM L25 (21 DUPLICATES REMOVED)

=> d 1-19 an ti so au

L26 ANSWER 1 OF 19 COPYRIGHT 1993 BIOSIS DUBPLICATE 1
AN 93:102377 BIOSIS
TI SELF-ASSEMBLY OF HUMAN PAPILLOMAVIRUS TYPE 1 CAPSIDS BY
   EXPRESSION OF THE L1 PROTEIN ALONE OR BY
   COEXPRESSION OF THE L1 AND L2 ***CAPSID*** PROTEINS.
SO J VIROL 67 (1). 1993. 315-322. CODEN: JOVIAM ISSN: 0022-538X
AU HAGENSEE M E; YAEGASHI N; GALLOWAY D A

L26 ANSWER 2 OF 19 COPYRIGHT 1993 BIOSIS DUBPLICATE 2
AN 93:102310 BIOSIS
TI PAPILLOMAVIRUS L1 MAJOR CAPSID PROTEIN
   SELF-ASSEMBLES INTO VIRUS-LIKE PARTICLES THAT ARE HIGHLY IMMUNOGENIC.
SO PROC NATL ACAD SCI U S A 89 (24). 1992. 12180-12184. CODEN: PNASA6
   ISSN: 0027-8424

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AU KIRNBAUER R; BOO ; CHENG N; LOWY D R; SCHI ER J T

L26 ANSWER 3 OF 19 COPYRIGHT 1993 BIOSIS DUPLICATE 3
AN 92:433005 BIOSIS
TI DEFINITION OF LINEAR ANTIGENIC REGIONS OF THE HPV16 L1 ***CAPSID***
PROTEIN USING SYNTHETIC VIRION-LIKE PARTICLES.
SO VIROLOGY 189 (2). 1992. 592-599. CODEN: VIRLAX ISSN: 0042-6822
AU ZHOU J; SUN X-Y; DAVIES H; CRAWFORD L; PARK D; FRAZER I H

L26 ANSWER 4 OF 19 COPYRIGHT 1993 BIOSIS DUPLICATE 4
AN 92:501971 BIOSIS
TI HPV-1 L1 PROTEIN EXPRESSED IN COS CELLS
DISPLAYS CONFORMATIONAL EPITOPES FOUND ON INTACT VIRIONS.
SO VIROLOGY 190 (1). 1992. 548-552. CODEN: VIRLAX ISSN: 0042-6822
AU GHIM S-J; JENSON A B; SCHLEGEL R

L26 ANSWER 5 OF 19 COPYRIGHT 1993 NLM DUPLICATE 5
AN 92364377 MEDLINE
TI Factors associated with detection of human papillomavirus
E4 and L1 proteins in condylomata acuminata.
SO J Infect Dis, (1992 Sep) 166 (3) 512-7
Journal code: IH3 ISSN: 0022-1899
AU Brown DR; Bryan JT; Rodriguez M; Katz BP

L26 ANSWER 6 OF 19 COPYRIGHT 1993 BIOSIS
AN 92:518271 BIOSIS
TI HUMAN PAPILLOMAVIRUS L1 PROTEIN IN
CONDYLOMATA ACUMINATA.
SO 32ND INTERSCIENCE CONFERENCE ON ANTIMICROBIAL AGENTS AND
CHEMOTHERAPY, ANAHEIM, CALIFORNIA, USA, OCTOBER 11-14, 1992. PROGRAM
ABSTR INTERSCI CONF ANTIMICROB AGENTS CHEMOTHERAPY 32 (0). 1992.
255. CODEN: POCHES
AU WOOLS K; BRYAN J; BROWN D

L26 ANSWER 7 OF 19 COPYRIGHT 1993 BIOSIS
AN 91:204291 BIOSIS
TI CHARACTERIZATION OF MURINE POLYCLONAL ANTISERA AND MONOCLONAL
ANTIBODIES GENERATED AGAINST INTACT AND DENATURED HUMAN
PAPILLOMAVIRUS TYPE 1 VIRIONS.
SO J VIROL 65 (3). 1991. 1578-1583. CODEN: JOVIAM ISSN: 0022-538X
AU YAEGASHI N; JENISON S A; VALENTINE J M; DUNN M; TAICHMAN L B; BAKER D
A; GALLOWAY D A

L26 ANSWER 8 OF 19 COPYRIGHT 1993 BIOSIS DUPLICATE 6
AN 92:103378 BIOSIS
TI IDENTIFICATION OF THE NUCLEAR LOCALIZATION SIGNAL OF HUMAN
PAPILLOMAVIRUS TYPE 16 L1 PROTEIN.
SO VIROLOGY 185 (2). 1991. 625-632. CODEN: VIRLAX ISSN: 0042-6822
AU ZHOU J; DOORBAR J; SUN X Y; CRAWFORD L V; MCLEAN C S; FRAZER I H

L26 ANSWER 9 OF 19 COPYRIGHT 1993 BIOSIS
AN 91:456420 BIOSIS
TI TYPE-SPECIFIC AND CROSS-REACTIVE EPITOPES IN HUMAN
PAPILLOMAVIRUS TYPE 16 CAPSID PROTEINS.
SO VIROLOGY 184 (1). 1991. 460-464. CODEN: VIRLAX ISSN: 0042-6822
AU BEISS B K; HEIMER E; FELIX A; BURK R D; RITTER D B; MALLON R G;
KADISH A S

L26 ANSWER 10 OF 19 COPYRIGHT 1993 NLM
AN 91134982 MEDLINE
TI The induction of cytotoxic T-lymphocyte precursor cells by
recombinant vaccinia virus expressing human papillomavirus

type 16 L1.

SO Virology, (1991 Mar) 181 (1) 203-10

Journal code: XEA ISSN: 0042-6822

AU Zhou JA; McIndoe A; Davies H; Sun XY; Crawford L

L26 ANSWER 11 OF 19 COPYRIGHT 1993 BIOSIS DUPLICATE 7

AN 91:295875 BIOSIS

TI HUMAN PAPILLOMAVIRUS TYPE 16 INFECTION OF THE CERVIX A
COMPARISON OF DIFFERING DNA DETECTION MODES AND THE USE OF MONOCLONAL
ANTIBODIES AGAINST THE MAJOR CAPSID PROTEIN.

SO GENITOURIN MED 67 (2). 1991. 87-91. CODEN: GEMEE2 ISSN: 0266-4348

AU LACEY C J N; WELLS M; MACDERMOTT R I J; GIBSON P E

L26 ANSWER 12 OF 19 COPYRIGHT 1993 BIOSIS DUPLICATE 8

AN 90:308849 BIOSIS

TI HUMAN PAPILLOMAVIRUS TYPE 1 PRODUCES REDUNDANT AS WELL AS
POLYCYSTRONIC MESSENGER RNA IN PLANTAR WARTS.

SO J VIROL 64 (6). 1990. 3144-3149. CODEN: JOVIAM ISSN: 0022-538X

AU PALERMO-DILTS D A; BROKER T S R; CHOW L T

L26 ANSWER 13 OF 19 COPYRIGHT 1993 BIOSIS

AN 91:49209 BIOSIS

TI DEFINITION OF MURINE T HELPER CELL DETERMINANTS IN THE MAJOR
CAPSID PROTEIN OF HUMAN PAPILLOMAVIRUS TYPE 16.

SO J GEN VIROL 71 (11). 1990. 2691-2698. CODEN: JGVIAI ISSN: 0022-1317

AU DAVIES D H; HILL C M; ROTHBARD J B; CHAIN B M

L26 ANSWER 14 OF 19 COPYRIGHT 1993 NLM

AN 91011369 MEDLINE

TI Increased antibody responses to human papillomavirus type
16 L1 protein expressed by recombinant vaccinia
virus lacking serine protease inhibitor genes.

SO J Gen Virol, (1990 Sep) 71 (Pt 9) 2185-90

Journal code: I9B ISSN: 0022-1317

AU Zhou J; Crawford L; McLean L; Sun XY; Stanley M; Almond N; Smith GL

L26 ANSWER 15 OF 19 COPYRIGHT 1993 BIOSIS DUPLICATE 9

AN 90:354554 BIOSIS

TI PRODUCTION AND CHARACTERIZATION OF A MONOCLONAL ANTIBODY TO HUMAN
PAPILLOMAVIRUS TYPE 16 USING RECOMBINANT VACCINIA VIRUS.

SO J CLIN PATHOL (LOND) 43 (6). 1990. 488-492. CODEN: JCPAAK ISSN:
0021-9746

AU MCLEAN C S; CHURCHER M J; MEINKE J; SMITH G L; HIGGINS G; STANLEY M;
MINSON A C

L26 ANSWER 16 OF 19 COPYRIGHT 1993 BIOSIS

AN 90:155714 BIOSIS

TI HUMORAL ASSAYS OF HUMAN SERA TO DISRUPTED AND NONDISRUPTED EPITOPES
OF HUMAN PAPILLOMAVIRUS TYPE 1.

SO VIROLOGY 174 (2). 1990. 388-398. CODEN: VIRLAX ISSN: 0042-6822

AU STEELE J C; GALLIMORE P H

L26 ANSWER 17 OF 19 COPYRIGHT 1993 BIOSIS DUPLICATE 10

AN 89:513896 BIOSIS

TI CHARACTERIZATION OF RARE HUMAN PAPILLOMAVIRUS TYPE 11
MESSENGER RNA SPECIES CODING FOR REGULATORY AND STRUCTURAL PROTEINS
USING THE POLYMERASE CHAIN REACTION.

SO VIROLOGY 172 (2). 1989. 489-497. CODEN: VIRLAX ISSN: 0042-6822

AU ROTENBERG M O; CHOW L T; BROKER T R

L26 ANSWER 18 OF 19 COPYRIGHT 1993 BIOSIS DUPLICATE 11

AN 89:161987 BIOSIS

* * * * *

```
=> s papillomavirus
L1      0 PAPILLOMAVIRIS
=>
=> s papillomavirus
L2      95 PAPILLOMAVIRUS
=> s "L!"
L3      426702 "L!"
        ("L")
```

```
=> s "L1"
L4      22478 "L1"
=> s l2 and l4
L5      13 L2 AND L4
=> d l5 1-13
```

1. 5,411,857, May 2, 1995, Probes for papillomaviruses and an in vitro diagnostic procedure for papilloma infections; Sylvie Beaudenon, et al., 435/5, 6; 536/23.72 [IMAGE AVAILABLE]

2. 5,401,627, Mar. 28, 1995, Antibodies to human **papillomavirus** latent proteins, diagnostic systems and methods; Joakim Dillner, et al., 435/5, 240.27; 436/518, 548; 530/387.9, 388.3, 389.4 [IMAGE AVAILABLE]

3. 5,364,758, Nov. 15, 1994, Primers and process for detecting human **papillomavirus** genotypes by PCR; Christophorus J. Meijer, et al., 435/5, 6, 91.2; 536/24.32, 24.33; 935/78 [IMAGE AVAILABLE]

4. 5,346,811, Sep. 13, 1994, Method and products for human **papillomavirus** detection; Ivan Galindo-Castro, et al., 435/5, 6; 530/387.1; 536/24.32 [IMAGE AVAILABLE]

5. 5,342,930, Aug. 30, 1994, Isolated DNA of human **papillomavirus** type 54(HPV54); Gerard Orth, et al., 536/23.72; 435/172.3, 320.1; 536/24.32 [IMAGE AVAILABLE]

6. 5,334,515, Aug. 2, 1994, Method for altering a nucleotide sequence; Ayoub Rashtchian, et al., 435/91.2, 91.41, 91.51, 172.3, 227 [IMAGE AVAILABLE]

7. 5,283,171, Feb. 1, 1994, Compositions for and detection of human **papillomavirus** by specific oligonucleotide polymerase primers using the polymerase chain reaction; M. Michele Manos, et al., 435/5, 6, 810; 436/501, 811; 536/23.1, 24.3, 24.31, 24.32, 24.33; 935/3, 20, 77, 78, 88 [IMAGE AVAILABLE]

8. 5,194,370, Mar. 16, 1993, Promoter ligation activated transcription amplification of nucleic acid sequences; Mark S. Berninger, et al., 435/6, 91.21; 436/94, 501; 935/77, 78 [IMAGE AVAILABLE]

9. 5,182,377, Jan. 26, 1993, Probes for detection of human **papillomavirus** M. Michele Manos, et al., 536/24.32; 435/5, 6; 436/501, 811; 536/24.33; 935/3, 20, 77, 78 [IMAGE AVAILABLE]

10. 5,180,806, Jan. 19, 1993, Polypeptides and compositions of human **papillomavirus** latent proteins, diagnostic systems and methods; Joakim Dillner, et al., 530/326, 324, 325 [IMAGE AVAILABLE]

=> s papillocavirus
L1 78 PAPILLOHAVIRUS

=> s "L1"
L2 20885 "L1"

=> s l1 and l2
L3 8 L1 AND L2

=> d l3 1-8

1. 5,334,515, Aug. 2, 1994, Method for altering a nucleotide sequence; Ayoub Rashtchian, et al., 435/91.2, 91.41, 91.51, 172.3, 227 [IMAGE AVAILABLE]

2. 5,283,171, Feb. 1, 1994, Compositions for and detection of human papillonavirus by specific oligonucleotide polymerase primers using the polymerase chain reaction; M. Michele Manos, et al., 435/5, 6, 810; 436/581, 811; 536/23.1, 24.3, 24.31, 24.32, 24.33; 935/3, 28, 77, 78, 88 [IMAGE AVAILABLE]

3. 5,194,378, Mar. 16, 1993, Promoter ligation activated transcription amplification of nucleic acid sequences; Mark S. Berninger, et al., 435/6, 91.21; 436/94, 581; 935/77, 78 [IMAGE AVAILABLE]

4. 5,182,377, Jan. 26, 1993, Probes for detection of human papillonavirus; M. Michele Manos, et al., 536/24.32; 435/5, 6; 436/581, 811; 536/24.33; 935/3, 28, 77, 78 [IMAGE AVAILABLE]

5. 5,180,885, Jan. 19, 1993, Polypeptides and compositions of human papillonavirus latent proteins, diagnostic systems and methods; Joakim Dillner, et al., 538/326, 324, 325 [IMAGE AVAILABLE]

6. 5,857,411, Oct. 15, 1991, Type-specific papillonavirus DNA sequences and peptides; Wayne D. Lancaster, et al., 435/6, 5; 436/581, 811; 536/23.72, 24.32; 935/78 [IMAGE AVAILABLE]

7. 4,886,741, Dec. 12, 1989, Use of volucre exclusion agents for the enhancement of in situ hybridization; Dennis E. Schwartz, 435/5, 6, 21, 810; 436/581; 935/77, 78 [IMAGE AVAILABLE]

8. 4,551,278, Nov. 5, 1985, DNA Fragments coding for polypeptides containing at least one antigenic determinant of the papillonavirus, particularly of the HPV 1a type and corresponding polypeptides; Olivier Danos, et al., 538/327, 329; 938/228, DIG.811 [IMAGE AVAILABLE]

FILE 'HUGHES' ENTERED AT 14:02:43 ON 16 AUG 94

FILE 'DB' ENTERED AT 14:20:25 ON 16 AUG 94

11 1767 S RABILLONOVIRUS/OI,BI
 12 1134 S HPU/OB,BI
 13 7012 S 30/OB,BI
 14 1572741 S 10/OB,BI
 15 894 S 011 OR 101 AND (LR OF LR)
 16 055051 S 010/OB,BI
 17 100 S L5 AND L1
 18 20 S L7 AND TYPE/OB,BI
 19 0 S 011 OR 101 AND (TYPE 10)/OB,BI
 20 0 S 011 OR 101 AND (TYPE 2)/OB,BI
 21 0 S 011 OR 101 AND (TYPE 3)/OB,BI

FILE 'MEDLINE' ENTERED AT 14:09:13 ON 16 AUG 94

112 0 S L11
 113 0 S L9

FILE 'DIOSIS' ENTERED AT 14:11:57 ON 16 AUG 94

114 0 S L12
 115 1 S L13

FILE 'DIOSAPS, ABI-INFORM' ENTERED AT 14:14:24 ON 16 AUG 94

> 100 113

FILE 'DIOSAPS'

'DB' IS NOT A VALID FIELD CODE

0 RABILLONOVIRUS/OR
 101 RABILLONOVIRUS/BI
 2 HPU/OB
 00 HPU/BI
 2 (TYPE 30)/OB
 04210 "TYPE"/BI
 422 "20"/BI
 0 (TYPE 30)/BI
 0 ("TYPE"(0) "20")/BI
 0 RABILLONOVIRUS/OR
 024 RABILLONOVIRUS/BI
 2 HPU/OB
 00 HPU/BI
 2 (TYPE 10)/OB
 04210 "TYPE"/BI
 04210 "10"/BI
 4 (TYPE 10)/BI
 0 ("TYPE"(0) "10")/BI
 0 L10 OR L12

FILE 'ABI-INFORM'

1 RABILLONOVIRUS/OR
 2 RABILLONOVIRUS/BI
 0 HPU/OB
 0 HPU/BI

17

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>>> C 18 30 all
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Enter "DISPLAY HISTORY" to see a list of the files in the current
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13 ANSWER IS OF ID CA COPYRIGHT 1994 AGO
14 ALL-RESIDE CA
15 Name: Sumpapilloraviracox ***** 26 (***** -80),
16 or ***** -Beralated ***** associated with skin pain.
17 Laura, Michael, Charles, Giovanni, Jekloraka, Stefania, Josh, Gerard
18 John, Paul, John, David, 77724, St.
19 P. Miller (1998), 1998, 1998
20 1998, 1998, 1998, 1998, 1998
21 1998

Relative alignment of the restriction endonuclease map of the 2
 1000 genome was 1.7 kb, as indicated for cloning and
 200 of phage 1.7 kb, so that the map of the 2000 of BSV-3
 300 can be compared with the BSV-1, both of which contain a
 400 1.7 kb region, there is a small gap, large deletion and exhibit
 500 genome divergence.
 600 Bovine papilloma virus
 700 Papillomavirus DNA
 800 (of bovine papilloma viruses types 3 and 4, colinearity of)
 900 Virus, animal
 1000 (Bovine papilloma, genomes of types 3 and 4 of, colinearity of)

CHAPER 2 OF 2 OF COPYRIGHT 1934 223
53.31483 00

Characterization of two types of human papillomaviruses in lesions of epidermodysplasia verruciformis.
Orléans, Gerard; Jabloncka, Stefania; Favre, Michel; Croissant, Odile; Jankovits-Thomassot, Marie; Rissa, Genevieve; Uitto Rech, Etienne. *Virale Cancers* Mon., Inst. Gustave-Roussy, Villejuif, Fr.
Prog. Natl. Acad. Sci. U. S. A. (1978), 75(3), 1537-41
GOST: PH0886: ISSN: 0037-0434

Journal
 54754

14-1 (Mammalian Pathological Biochemistry)
Enzymes: 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29, 30, 31, 32, 33, 34, 35, 36, 37, 38, 39, 40, 41, 42, 43, 44, 45, 46, 47, 48, 49, 50, 51, 52, 53, 54, 55, 56, 57, 58, 59, 60, 61, 62, 63, 64, 65, 66, 67, 68, 69, 70, 71, 72, 73, 74, 75, 76, 77, 78, 79, 80, 81, 82, 83, 84, 85, 86, 87, 88, 89, 90, 91, 92, 93, 94, 95, 96, 97, 98, 99, 100, 101, 102, 103, 104, 105, 106, 107, 108, 109, 110, 111, 112, 113, 114, 115, 116, 117, 118, 119, 120, 121, 122, 123, 124, 125, 126, 127, 128, 129, 130, 131, 132, 133, 134, 135, 136, 137, 138, 139, 140, 141, 142, 143, 144, 145, 146, 147, 148, 149, 150, 151, 152, 153, 154, 155, 156, 157, 158, 159, 160, 161, 162, 163, 164, 165, 166, 167, 168, 169, 170, 171, 172, 173, 174, 175, 176, 177, 178, 179, 180, 181, 182, 183, 184, 185, 186, 187, 188, 189, 190, 191, 192, 193, 194, 195, 196, 197, 198, 199, 200, 201, 202, 203, 204, 205, 206, 207, 208, 209, 210, 211, 212, 213, 214, 215, 216, 217, 218, 219, 220, 221, 222, 223, 224, 225, 226, 227, 228, 229, 230, 231, 232, 233, 234, 235, 236, 237, 238, 239, 240, 241, 242, 243, 244, 245, 246, 247, 248, 249, 250, 251, 252, 253, 254, 255, 256, 257, 258, 259, 260, 261, 262, 263, 264, 265, 266, 267, 268, 269, 270, 271, 272, 273, 274, 275, 276, 277, 278, 279, 280, 281, 282, 283, 284, 285, 286, 287, 288, 289, 290, 291, 292, 293, 294, 295, 296, 297, 298, 299, 300, 301, 302, 303, 304, 305, 306, 307, 308, 309, 310, 311, 312, 313, 314, 315, 316, 317, 318, 319, 320, 321, 322, 323, 324, 325, 326, 327, 328, 329, 330, 331, 332, 333, 334, 335, 336, 337, 338, 339, 340, 341, 342, 343, 344, 345, 346, 347, 348, 349, 350, 351, 352, 353, 354, 355, 356, 357, 358, 359, 360, 361, 362, 363, 364, 365, 366, 367, 368, 369, 370, 371, 372, 373, 374, 375, 376, 377, 378, 379, 380, 381, 382, 383, 384, 385, 386, 387, 388, 389, 390, 391, 392, 393, 394, 395, 396, 397, 398, 399, 400, 401, 402, 403, 404, 405, 406, 407, 408, 409, 410, 411, 412, 413, 414, 415, 416, 417, 418, 419, 420, 421, 422, 423, 424, 425, 426, 427, 428, 429, 430, 431, 432, 433, 434, 435, 436, 437, 438, 439, 440, 441, 442, 443, 444, 445, 446, 447, 448, 449, 450, 451, 452, 453, 454, 455, 456, 457, 458, 459, 460, 461, 462, 463, 464, 465, 466, 467, 468, 469, 470, 471, 472, 473, 474, 475, 476, 477, 478, 479, 480, 481, 482, 483, 484, 485, 486, 487, 488, 489, 490, 491, 492, 493, 494, 495, 496, 497, 498, 499, 500, 501, 502, 503, 504, 505, 506, 507, 508, 509, 510, 511, 512, 513, 514, 515, 516, 517, 518, 519, 520, 521, 522, 523, 524, 525, 526, 527, 528, 529, 530, 531, 532, 533, 534, 535, 536, 537, 538, 539, 540, 541, 542, 543, 544, 545, 546, 547, 548, 549, 550, 551, 552, 553, 554, 555, 556, 557, 558, 559, 560, 561, 562, 563, 564, 565, 566, 567, 568, 569, 570, 571, 572, 573, 574, 575, 576, 577, 578, 579, 580, 581, 582, 583, 584, 585, 586, 587, 588, 589, 590, 591, 592, 593, 594, 595, 596, 597, 598, 599, 600, 601, 602, 603, 604, 605, 606, 607, 608, 609, 610, 611, 612, 613, 614, 615, 616, 617, 618, 619, 620, 621, 622, 623, 624, 625, 626, 627, 628, 629, 630, 631, 632, 633, 634, 635, 636, 637, 638, 639, 640, 641, 642, 643, 644, 645, 646, 647, 648, 649, 650, 651, 652, 653, 654, 655, 656, 657, 658, 659, 660, 661, 662, 663, 664, 665, 666, 667, 668, 669, 670, 671, 672, 673, 674, 675, 676, 677, 678, 679, 680, 681, 682, 683, 684, 685, 686, 687, 688, 689, 690, 691, 692, 693, 694, 695, 696, 697, 698, 699, 700, 701, 702, 703, 704, 705, 706, 707, 708, 709, 710, 711, 712, 713, 714, 715, 716, 717, 718, 719, 720, 721, 722, 723, 724, 725, 726, 727, 728, 729, 730, 731, 732, 733, 734, 735, 736, 737, 738, 739, 740, 741, 742, 743, 744, 745, 746, 747, 748, 749, 750, 751, 752, 753, 754, 755, 756, 757, 758, 759, 760, 761, 762, 763, 764, 765, 766, 767, 768, 769, 770, 771, 772, 773, 774, 775, 776, 777, 778, 779, 780, 781, 782, 783, 784, 785, 786, 787, 788, 789, 790, 791, 792, 793, 794, 795, 796, 797, 798, 799, 800, 801, 802, 803, 804, 805, 806, 807, 808, 809, 810, 811, 812, 813, 814, 815, 816, 817, 818, 819, 820, 821, 822, 823, 824, 825, 826, 827, 828, 829, 830, 831, 832, 833, 834, 835, 836, 837, 838, 839, 840, 841, 842, 843, 844, 845, 846, 847, 848, 849,

Human papillomaviruses (HPVs) found in lesions of 11 patients suffering from epidermodysplasia verruciformis were compared to HPV-viruses type 1 (HPV-1) and type 2 (HPV-2) previously characterized in plantar and common warts, resp. Complementary RNAs (cRNAs) to HPV-1, HPV-2, and viruses from 2 patients with epidermodysplasia verruciformis (J.D. HPV-3 and J.K. HPV-4) were used in cRNA-DNA filter hybridization experiments. No sequence homology was detected between HPV-1 or HPV-2 and HPV-3 DNAs and DNAs obtained from the 11 epidermodysplasia verruciformis HPV-3 isolates. Furthermore, with J.D. and J.K. HPV-4 cRNAs, epidermodysplasia verruciformis HPV-3 DNAs fall into 2 groups showing little, if any, sequence homology. A lower extent of similarity was found for the DNAs of some isolates showing a genetic heterogeneity within each of the groups. Almost no antigenic cross-reactivity was detected by immunodiffusion and indirect immunofluorescence tests, either between epidermodysplasia verruciformis HPV-3 and HPV-1 or HPV-2 or between J.D. and J.K. HPVs. Viruses belonging to the same group have common antigenic properties, but antigenic differences were also shown for the viruses having only partial DNA sequence homology. Viruses related to J.D. HPV-3 were preferentially found with flat wart-like lesions of epidermodysplasia verruciformis and were further found in the lesions of 5 patients bearing multiple flat warts. Viruses related to J.K. HPV-4 were found in multiple distinct lesions (red spots) present in some patients with epidermodysplasia verruciformis. Thus, it is proposed to distinguish 5 other types of HPV designated provisionally as HPV-5 (HPV-5), HPV-6 (HPV-6), HPV-7 (HPV-7), HPV-8 (HPV-8) and type 4 (HPV-4), with J.D. and J.K. HPVs as prototypes, resp. Malignant conversion of some epidermodysplasia verruciformis lesions is more frequently associated with HPV-4 than with HPV-3 infection.

1. *Epidermoplastus verreauxi* (Lacaze) (Fig. 1)
 2. *Epidermoplastus verreauxi* (Lacaze) (Fig. 2)
 3. *Epidermoplastus verreauxi* (Lacaze) (Fig. 3)
 4. *Epidermoplastus verreauxi* (Lacaze) (Fig. 4)
 5. *Epidermoplastus verreauxi* (Lacaze) (Fig. 5)
 6. *Epidermoplastus verreauxi* (Lacaze) (Fig. 6)
 7. *Epidermoplastus verreauxi* (Lacaze) (Fig. 7)
 8. *Epidermoplastus verreauxi* (Lacaze) (Fig. 8)
 9. *Epidermoplastus verreauxi* (Lacaze) (Fig. 9)
 10. *Epidermoplastus verreauxi* (Lacaze) (Fig. 10)

From this, we can see that the α value is 0.05, and the β value is 0.05. The α value is the probability of rejecting the null hypothesis when it is true, and the β value is the probability of accepting the null hypothesis when it is false. The α value is also known as the Type I error, and the β value is also known as the Type II error. The α value is usually set at 0.05, and the β value is usually set at 0.05. The α value is the probability of rejecting the null hypothesis when it is true, and the β value is the probability of accepting the null hypothesis when it is false. The α value is also known as the Type I error, and the β value is also known as the Type II error.

Human papillomaviruses associated with epidermodysplasia verruciformis (HVPV-11). Molecular cloning and biochemical characterization of human papillomaviruses associated with epidermodysplasia verruciformis (HVPV-11).

Koussis, Dina, Iakobson, Stefania, Favre, Michel, Orth, Gerard, Suter, Sabine, Inat, Sabir, Suter, Sabine, Paris, 75015, France, J. Virol. (1993), 67(12), 342-51
 000000 000000 000000 000000 000000 000000 000000 000000

Journal

Epidermis

3-2 (Biochemical Genetics)

Genetic, cross-referenced to 14

The DNAs of 4 human papillomaviruses (HPVs) that were found in the benign lesions of 3 patients suffering from epidermodysplasia verruciformis were characterized. The flat, wart-like lesions and the nodular lesions of patient 1 contained 2 viruses, HPV-3a and HPV-9, resp., whose genomes had previously been only partially characterized. The flat, wart-like lesions of patient 2 and the nodular lesions of patient 3 each contained a virus previously considered as belonging to types 3 and 5, resp. These viruses are different from all the HPV types so far characterized; they have tentatively been named HPV-10 and HPV-12. The HPV-3a, HPV-9, and HPV-12 DNAs and the 2 SalI fragments of HPV-10 DNA (54.1 and 5.6% of the genome length) were cloned in Escherichia coli after having been inserted in plasmid pBR322. The cloned HPV genomes have similar sizes (approx. 7700 base pairs), but their guanine-plus-cytosine contents differ from 41.8% for HPV-12 DNA to 45.5% for HPV-3a DNA. The study of the sensitivity of the 4 HPV DNAs to 14 restriction endonucleases permitted the construction of cleavage maps. Evidence for conserved restriction sites was found only for the HPV-3a and HPV-10 genome, since 5 of the 21 restriction sites localized in the HPV-3a DNA were also to be present in the HPV-10 DNA. Hybridization experiments, performed in liq. phase at 60°C, showed a 50% sequence homol. between HPV-3a and HPV-10 DNAs, 17-23% sequence homol. among HPV-3, HPV-9, and HPV-12 DNAs, almost no sequence homol. between the HPV-3a or HPV-10 DNA and the other HPV DNAs, and a weak homol. between HPV-9 DNA and HPV-8 or HPV-12 DNA. Blot hybridization experiments showed no sequence homol. between the HPV-3a, HPV-9, and HPV-12 DNAs and the DNAs of the HPVs assoc. with skin warts (HPV-1, HPV-2, HPV-4, and HPV-7) or with mucocutaneous and mucous membrane lesions (HPV-6 and HPV-11a, resp.). One exception was a weak sequence homol. between the HPV-2 prototype and HPV-3a or HPV-10 DNA. Thus, at least the following 6 HPV types are assoc. with epidermodysplasia verruciformis: HPV-3a and HPV-10, which are assoc. with flat warts in the general population; and HPV-9, HPV-8, HPV-5, and HPV-12, which are assoc. specifically with epidermodysplasia verruciformis.

Human papillomaviruses associated with epidermodysplasia verruciformis

Human papillomavirus associated with epidermodysplasia verruciformis

Molecular cloning

(of DNA, of human papillomaviruses assoc. with epidermodysplasia verruciformis)

Deoxyribonucleic acids

(of human papillomaviruses assoc. with epidermodysplasia verruciformis, characterization of)

Skin, disease or disorder

Epidermodysplasia verruciformis, DNA of human papillomaviruses associated with, cloning and characterization of

Virus, animal

(papilloma, DNA of, assoc. with epidermodysplasia verruciformis, cloning and characterization of)